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(54) Title: METHODS OF DIAGNOSIS OF PROSTATE CANCER, COMPOSITIONS AND METHODS OF SCREENING FOR MODULATORS OF PROSTATE CANCER

2 (57) Abstract: Described herein are genes whose expression are up-regulated or down-regulated in prostate cancer. Also described are such genes whose expression is further up-regulated or down-regulated in drug-resistant prostate cancer cells. Related methods and compositions that can be used for diagnosis and treatment of prostate cancer are disclosed. Also described herein are methods that can be used to identify modulators of prostate cancer.

# METHODS OF DIAGNOSIS OF PROSTATE CANCER, COMPOSITIONS AND METHODS OF SCREENING FOR MODULATORS OF PROSTATE CANCER

#### CROSS-REFERENCES TO RELATED APPLICATIONS

This application claims priority from the following applications: USSN 09/687,576 filed October 13, 2000, USSN 60/276,791 filed March 16, 2001; USSN 60/288,589, filed May 4, 2001; USSN 09/733,742, filed December 8, 2000; USSN 09/733,288, filed December 8, 2000; USSN 09/847,046, filed April 30, 2001; USSN 60/276,888, filed March 16, 2001; USSN 60/286,214, filed April 24, 2001; USSN 60/281,922, filed April 6, 2001; USSN 60/263,957, filed January 24, 2001, which are incorporated herein by reference in their entirety.

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#### FIELD OF THE INVENTION

The invention relates to the identification of nucleic acid and protein expression profiles and nucleic acids, products, and antibodies thereto that are involved in prostate cancer; and to the use of such expression profiles and compositions in the diagnosis, prognosis and therapy of prostate cancer. The invention further relates to methods for identifying and using agents and/or targets that inhibit prostate cancer.

#### BACKGROUND OF THE INVENTION

Prostate cancer is the most commonly diagnosed internal malignancy and second most common cause of cancer death in men in the U.S., resulting in approximately 40,000 deaths each year (Landis et al., CA Cancer J. Clin. 48.6-29 (1998); Greenlee et al., CA Cancer J. Clin. 50(1):7-13 (2000)), and incidence of prostate cancer has been increasing rapidly over the past 20 years in many parts of the world (Nakata et al., Int. J. Urol. 7(7):254-257 (2000); Majeed et al., BJU Int. 85(9):1058-1062 (2000)). It develops as the

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result of a pathologic transformation of normal prostate cells. In tumorigenesis, the cancer cell undergoes initiation, proliferation and loss of contact inhibition, culminating in invasion of surrounding tissue and, ultimately, metastasis.

Deaths from prostate cancer are a result of metastasis of a prostate tumor.

Therefore, early detection of the development of prostate cancer is critical in reducing mortality from this disease. Measuring levels of prostate-specific antigen (PSA) has become a very common method for early detection and screening, and may have contributed to the slight decrease in the mortality rate from prostate cancer in recent years (Nowroozi et al., Cancer Control 5(6):522-531 (1998)). However, many cases are not diagnosed until the disease has progressed to an advanced stage.

Treatments such as surgery (prostatectomy), radiation therapy, and cryotherapy are potentially curative when the cancer remains localized to the prostate. Therefore, early detection of prostate cancer is important for a positive prognosis for treatment. Systemic treatment for metastatic prostate cancer is limited to hormone therapy and chemotherapy. Chemical or surgical castration has been the primary treatment for symptomatic metastatic prostate cancer for over 50 years. This testicular androgen deprivation therapy usually results in stabilization or regression of the disease (in 80% of patients), but progression of metastatic prostate cancer eventually develops (Panvichian et al., Cancer Control 3(6):493-500 (1996)). Metastatic disease is currently considered incurable, and the primary goals of treatment are to prolong survival and improve quality of life (Rago, Cancer Control 5(6):513-521 (1998)).

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Thus, methods that can be used for diagnosis and prognosis of prostate cancer and effective treatment of prostate cancer, and including particularly metastatic prostate cancer, would be desirable. Accordingly, provided herein are methods that can be used in diagnosis and prognosis of prostate cancer. Further provided are methods that can be used to screen candidate bioactive agents for the ability to modulate, e.g., treat, prostate cancer. Additionally, provided herein are molecular targets and compositions for therapeutic intervention in prostate cancer and other cancers.

# SUMMARY OF THE INVENTION

The present invention therefore provides nucleotide sequences of genes that
are up- and down-regulated in prostate cancer cells. Such genes are useful for diagnostic
purposes, and also as targets for screening for therapeutic compounds that modulate prostate
cancer, such as hormones or antibodies. Other aspects of the invention will become apparent
to the skilled artisan by the following description of the invention.

In one aspect, the present invention provides a method of detecting a prostate cancer-associated transcript in a cell from a patient, the method comprising contacting a biological sample from the patient with a polynucleotide that selectively hybridizes to a sequence at least 80% identical to a sequence as shown in Tables 1-16.

In one embodiment, the present invention provides a method of determining the level of a prostate cancer associated transcript in a cell from a patient.

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label.

In one embodiment, the present invention provides a method of detecting a prostate cancer-associated transcript in a cell from a patient, the method comprising

contacting a biological sample from the patient with a polynucleotide that selectively hybridizes to a sequence at least 80% identical to a sequence as shown in Tables 1-16.

In one embodiment, the polynucleotide selectively hybridizes to a sequence at least 95% identical to a sequence as shown in Tables 1-16. In another embodiment, the polynucleotide comprises a sequence as shown in Tables 1-16.

In one embodiment, the biological sample is a tissue sample. In another embodiment, the biological sample comprises isolated nucleic acids, e.g., mRNA.

In one embodiment, the polynucleotide is labeled, e.g., with a fluorescent

In one embodiment, the polynucleotide is immobilized on a solid surface.

In one embodiment, the patient is undergoing a therapeutic regimen to treat prostate cancer. In another embodiment, the patient is suspected of having metastatic prostate cancer.

In one embodiment, the patient is a human.

In one embodiment, the patient is suspected of having a taxol-resistant cancer.

In one embodiment, the prostate cancer associated transcript is mRNA.

In one embodiment, the method further comprises the step of amplifying nucleic acids before the step of contacting the biological sample with the polynucleotide.

In another aspect, the present invention provides a method of monitoring the efficacy of a therapeutic treatment of prostate cancer, the method comprising the steps of: (i) providing a biological sample from a patient undergoing the therapeutic treatment; and (ii) 5 determining the level of a prostate cancer-associated transcript in the biological sample by contacting the biological sample with a polynucleotide that selectively hybridizes to a sequence at least 80% identical to a sequence as shown in Tables 1-16, thereby monitoring the efficacy of the therapy. In a further embodiment, the patient has metastatic prostate cancer. In a further embodiment, the patient has a drug resistant (e.g., taxol resistant) form of prostate cancer.

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In one embodiment, the method further comprises the step of: (iii) comparing the level of the prostate cancer-associated transcript to a level of the prostate cancerassociated transcript in a biological sample from the patient prior to, or earlier in, the therapeutic treatment.

Additionally, provided herein is a method of evaluating the effect of a candidate prostate cancer drug comprising administering the drug to a patient and removing a cell sample from the patient. The expression profile of the cell is then determined. This method may further comprise comparing the expression profile to an expression profile of a healthy individual. In a preferred embodiment, said expression profile includes a gene of Tables 1-16.

In one aspect, the present invention provides an isolated nucleic acid molecule consisting of a polynucleotide sequence as shown in Tables 1-16.

In one embodiment, an expression vector or cell comprises the isolated nucleic

In one aspect, the present invention provides an isolated polypeptide which is encoded by a nucleic acid molecule having polynucleotide sequence as shown in Tables 1-16. In another aspect, the present invention provides an antibody that specifically binds to an isolated polypeptide which is encoded by a nucleic acid molecule having polynucleotide sequence as shown in Tables 1-16.

In one embodiment, the antibody is conjugated to an effector component, e.g., a fluorescent label, a radioisotope or a cytotoxic chemical.

In one embodiment, the antibody is an antibody fragment. In another embodiment, the antibody is humanized.

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In one aspect, the present invention provides a method of detecting a prostate cancer cell in a biological sample from a patient, the method comprising contacting the biological sample with an antibody as described herein.

In another aspect, the present invention provides a method of detecting antibodies specific to prostate cancer in a patient, the method comprising contacting a biological sample from the patient with a polypeptide encoded by a nucleic acid comprising a sequence from Tables 1-16.

In another aspect, the present invention provides a method for identifying a compound that modulates a prostate cancer-associated polypeptide, the method comprising the steps of: (i) contacting the compound with a prostate cancer-associated polypeptide, the polypeptide encoded by a polynucleotide that selectively hybridizes to a sequence at least 80% identical to a sequence as shown in Tables 1-16; and (ii) determining the functional effect of the compound upon the polypeptide.

In one embodiment, the functional effect is a physical effect, an enzymatic effect, or a chemical effect.

In one embodiment, the polypeptide is expressed in a eukaryotic host cell or cell membrane. In another embodiment, the polypeptide is recombinant.

In one embodiment, the functional effect is determined by measuring ligand binding to the polypeptide.

In another aspect, the present invention provides a method of inhibiting proliferation of a prostate cancer-associated cell to treat prostate cancer in a patient, the method comprising the step of administering to the subject a therapeutically effective amount of a compound identified as described herein.

In one embodiment, the compound is an antibody.

In another aspect, the present invention provides a drug screening assay

30 comprising the steps of: (i) administering a test compound to a mammal having prostate

cancer or to a cell sample isolated therefrom; (ii) comparing the level of gene expression of a

polynucleotide that selectively hybridizes to a sequence at least 80% identical to a sequence as shown in Tables 1-16 in a treated cell or mammal with the level of gene expression of the polynucleotide in a control cell sample or mammal, wherein a test compound that modulates the level of expression of the polynucleotide is a candidate for the treatment of prostate cancer.

In one embodiment, the control is a mammal with prostate cancer or a cell sample therefrom that has not been treated with the test compound. In another embodiment, the control is a normal cell or mammal.

In one embodiment, the test compound is administered in varying amounts or concentrations. In another embodiment, the test compound is administered for varying time periods. In another embodiment, the comparison can occur after addition or removal of the drug candidate.

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In one embodiment, the levels of a plurality of polynucleotides that selectively hybridize to a sequence at least 80% identical to a sequence as shown in Tables 1-16 are individually compared to their respective levels in a control cell sample or mammal. In a preferred embodiment the plurality of polynucleotides is from three to ten.

In another aspect, the present invention provides a method for treating a mammal having prostate cancer comprising administering a compound identified by the assay described herein.

In another aspect, the present invention provides a pharmaceutical composition for treating a mammal having prostate cancer, the composition comprising a compound identified by the assay described herein and a physiologically acceptable excipient.

In one aspect, the present invention provides a method of screening drug candidates by providing a cell expressing a gene that is up- and down-regulated as in a prostate cancer. In one embodiment, a gene is selected from Tables 1-16. The method further includes adding a drug candidate to the cell and determining the effect of the drug candidate on the expression of the expression profile gene.

In one embodiment, the method of screening drug candidates includes comparing the level of expression in the absence of the drug candidate to the level of expression in the presence of the drug candidate, wherein the concentration of the drug

candidate can vary when present, and wherein the comparison can occur after addition or removal of the drug candidate. In a preferred embodiment, the cell expresses at least two expression profile genes. The profile genes may show an increase or decrease.

Also provided is a method of evaluating the effect of a candidate prostate cancer drug comprising administering the drug to a transgenic animal expressing or over-expressing the prostate cancer modulatory protein, or an animal lacking the prostate cancer modulatory protein, for example as a result of a gene knockout.

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Moreover, provided herein is a biochip comprising one or more nucleic acid segments of Tables 1-16, wherein the biochip comprises fewer than 1000 nucleic acid probes. Preferably, at least two nucleic acid segments are included. More preferably, at least three nucleic acid segments are included.

Furthermore, a method of diagnosing a disorder associated with prostate cancer is provided. The method comprises determining the expression of a gene of Tables 1-16, in a first tissue type of a first individual, and comparing the distribution to the expression of the gene from a second normal tissue type from the first individual or a second unaffected individual. A difference in the expression indicates that the first individual has a disorder associated with prostate cancer.

In a further embodiment, the biochip also includes a polynucleotide sequence of a gene that is not up- and down-regulated in prostate cancer.

In one embodiment a method for screening for a bioactive agent capable of interfering with the binding of a prostate cancer modulatory protein) or a fragment thereof and an antibody which binds to said prostate cancer modulatory protein or fragment thereof. In a preferred embodiment, the method comprises combining a prostate cancer modulatory protein or fragment thereof, a candidate bioactive agent and an antibody which binds to said prostate cancer modulatory protein or fragment thereof. The method further includes determining the binding of said prostate cancer modulatory protein or fragment thereof and said antibody. Wherein there is a change in binding, an agent is identified as an interfering agent. The interfering agent can be an agonist or an antagonist. Preferably, the agent inhibits prostate cancer.

Also provided herein are methods of eliciting an immune response in an individual. In one embodiment a method provided herein comprises administering to an

individual a composition comprising a prostate cancer modulating protein, or a fragment thereof. In another embodiment, the protein is encoded by a nucleic acid selected from those of Tables 1-16.

Further provided herein are compositions capable of eliciting an immune response in an individual. In one embodiment, a composition provided herein comprises a prostate cancer modulating protein, preferably encoded by a nucleic acid of Tables 1-16, or a fragment thereof, and a pharmaceutically acceptable carrier. In another embodiment, said composition comprises a nucleic acid comprising a sequence encoding a prostate cancer modulating protein, preferably selected from the nucleic acids of Tables 1-16, and a 10 pharmaceutically acceptable carrier.

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Also provided are methods of neutralizing the effect of a prostate cancer protein, or a fragment thereof, comprising contacting an agent specific for said protein with said protein in an amount sufficient to effect neutralization. In another embodiment, the protein is encoded by a nucleic acid selected from those of Tables 1-16.

In another aspect of the invention, a method of treating an individual for prostate cancer is provided. In one embodiment, the method comprises administering to said individual an inhibitor of a prostate cancer modulating protein. In another embodiment, the method comprises administering to a patient having prostate cancer an antibody to a prostate cancer modulating protein conjugated to a therapeutic moiety. Such a therapeutic moiety can be a cytotoxic agent or a radioisotope.

# DETAILED DESCRIPTION OF THE INVENTION

In accordance with the objects outlined above, the present invention provides novel methods for diagnosis and prognosis evaluation for prostate cancer (PC), including metastatic prostate cancer, as well as methods for screening for compositions which modulate prostate cancer. Also provided are methods for treating prostate cancer.

In addition to the other nucleic acid and peptide sequences, the present invention also relates to the identification of PAA2 as a gene that is highly over expressed in prostate cancer patient tissues. PAA2 sequence is identical to the zinc transporter ZNT4. Results presented herein demonstrate that PAA2/ZNT4 is highly expressed in prostate cancer cells. The prostate gland is unique in that it has the highest capacity of any organ in the body

to accumulate zinc. Zinc uptake is regulated by prolactin and testosterone, which induce the expression of a member of the ZIP family of zinc transporters (Costello et al., 1999, J. Biol. Chem. 274:17499-17504). Zinc accumulation in the prostate functions to inhibit citrate oxidation, which results in a decrease in cellular ATP production (Costello and Franklin, 1998. Prostate 35:285-296). Cancer cells are more sensitive to decreased ATP production and have evolved to prevent zinc accumulation. Without wishing to be bound by theory, the up-regulation of ZNT4 in prostate cancer cells may result in protection of the cells from high zinc levels by its ability to pump accumulated zinc out of the cells.

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The present invention also relates to nucleic acid sequencess encoding PBH1. PBH1 is related to human TRPC7 (transient receptor potential-related channels, NP\_003298), a putative calcium channel highly expressed in brain (Nagamine et al., Genomics 54:124-131 (1998)). Trp is related to melastatin, a gene down-regulated in metastatic melanomas (Duncan et al., Cancer Res. 58:1515-1520 (1998)), and MTR1, a gene locallized to within the Beckwith-Wiedemann syndrome/Wilm's tumor susceptability region (Prawitt et al., Hum. Mol. Genet, 9:203-216 (2000)). Without wishing to be bound by theory, it is believed that PBH1 functions as a calcium channel.

As a calcium channel, PBH1 is an ideal target for a small molecule therapeutic, or a therapeutic antibody that disrupts channel function. CD20, the target of Rituximab in non-Hodgekin's lymphoma (Maloney et al., Blood 90:2188-2195 (1997); Leget and Czuczman, Curr. Opin. Oncol. 10:548-551 (1998)), is a plasma membrane calcium channel expressed in B cells (Tedder and Engel, Immunol. Today 15:450-454 (1994)). Similarly, a small molecule, or antibody that inhibits or alters a calcium signal mediated by PBH1, will result in the death of prostate cancer cells.

PBH1, and other genes of the invention, are also be useful as targets for cytotoxic T-lymphocytes. Genes that are tumor specific, or that are expressed in immuneprivileged organs, are currently being used as potential vaccine targets (Van den Eynde and Boon, Int. J. Clin. Lab. Res. 27:81-86 (1997)). The expression pattern of PBH1 indicates that it is an ideal target for cytotoxic T-lymphocytes. Thus, therapies that utilize PBH1-specific cytotoxic T-lymphocytes to induce prostate cancer cell death are also provided by this invention. See, e.g., U.S. Patent No. 6.051,227 and WO 00/32231, the disclosures of which are herein incorporated by reference.

The present invention is also related to the identification of PAA3 as a gene that is important in the modulation of prostate cancer and or breast cancer.

Tables 1-16 provide unigene cluster identification numbers, exemplar
accession numbers, or genomic nucleotide position numbers for the nucleotide sequence of
genes that exhibit increased or decreased expression in prostate cancer samples.

#### Definitions

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The term "prostate cancer protein" or "prostate cancer polynucleotide" or "prostate cancer-associated transcript" refers to nucleic acid and polypeptide polymorphic variants, alleles, mutants, and interspecies homologues that: (1) have a nucleotide sequence that has greater than about 60% nucleotide sequence identity, 65%, 70%, 75%, 80%, 85%, 90%, preferably 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98% or 99% or greater nucleotide sequence identity, preferably over a region of over a region of at least about 25, 50, 100, 200, 500, 1000, or more nucleotides, to a nucleotide sequence of or associated with a unigene cluster of Tables 1-16; (2) bind to antibodies, e.g., polyclonal antibodies, raised against an immunogen comprising an amino acid sequence encoded by a nucleotide sequence of or associated with a unigene cluster of Tables 1-16, and conservatively modified variants thereof; (3) specifically hybridize under stringent hybridization conditions to a nucleic acid sequence, or the complement thereof of Tables 1-16 and conservatively modified variants thereof or (4) have an amino acid sequence that has greater than about 60% amino acid sequence identity, 65%, 70%, 75%, 80%, 85%, 90%, preferably 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98% or 99% or greater amino sequence identity, preferably over a region of over a region of at least about 25, 50, 100, 200, 500, 1000, or more amino acid, to an amino acid sequence encoded by a nucleotide sequence of or associated with a unigene cluster of Tables 1-16. A polynucleotide or polypeptide sequence is typically from a mammal including, but not limited to, primate, e.g., human; rodent, e.g., rat, mouse, hamster; cow, pig, horse, sheep, or other mammal. A "prostate cancer polypeptide" and a "prostate cancer polynucleotide," include both naturally occurring or recombinant forms.

A "full length" prostate cancer protein or nucleic acid refers to a prostate cancer polypeptide or polynucleotide sequence, or a variant thereof, that contains all of the elements normally contained in one or more naturally occurring, wild type prostate cancer

polynucleotide or polypeptide sequences. For example, a full length prostate cancer nucleic acid will typically comprise all of the exons that encode for the full length, naturally ocurring protein. The "full length" may be prior to, or after, various stages of post-translation processing or splicing, including alternative splicing.

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"Biological sample" as used herein is a sample of biological tissue or fluid that contains nucleic acids or polypeptides, e.g., of a prostate cancer protein, polynucleotide or transcript. Such samples include, but are not limited to, tissue isolated from primates, e.g., humans, or rodents, e.g., mice, and rats. Biological samples may also include sections of tissues such as biopsy and autopsy samples, frozen sections taken for histologic purposes, blood, plasma, serum, sputum, stool, tears, mucus, hair, skin, etc. Biological samples also include explants and primary and/or transformed cell cultures derived from patient tissues. A biological sample is typically obtained from a cukaryotic organism, most preferably a mammal such as a primate e.g., chimpanzee or human; cow; dog; cat; a rodent, e.g., guinea pig, rat, mouse; rabbit; or a bird; reptile; or fish.

"Providing a biological sample" means to obtain a biological sample for use in methods described in this invention. Most often, this will be done by removing a sample of cells from an animal, but can also be accomplished by using previously isolated cells (e.g., isolated by another person, at another time, and/or for another purpose), or by performing the methods of the invention in vivo. Archival tissues, having treatment or outcome history, will be particularly useful.

The terms "identical" or percent "identity," in the context of two or more nucleic acids or polypeptide sequences, refer to two or more sequences or subsequences that are the same or have a specified percentage of amino acid residues or nucleotides that are the same (i.e., about 60% identity, preferably 70%, 75%, 80%, 85%, 90%, 91%, 92%, 93%, 94%, 95%, 96%, 97%, 98%, 99%, or higher identity over a specified region, when compared and aligned for maximum correspondence over a comparison window or designated region) as measured using a BLAST or BLAST 2.0 sequence comparison algorithms with default parameters described below, or by manual alignment and visual inspection (see, e.g., NCBI web site http://www.ncbi.nlm.nih.gov/BLAST/ or the like). Such sequences are then said to be "substantially identical." This definition also refers to, or may be applied to, the compliment of a test sequence. The definition also includes sequences that have deletions

and/or additions, as well as those that have substitutions, as well as naturally occurring, e.g., polymorphic or allelic variants, and man-made variants. As described below, the preferred algorithms can account for gaps and the like. Preferably, identity exists over a region that is at least about 25 amino acids or nucleotides in length, or more preferably over a region that is 50-100 amino acids or nucleotides in length.

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For sequence comparison, typically one sequence acts as a reference sequence, to which test sequences are compared. When using a sequence comparison algorithm, test and reference sequences are entered into a computer, subsequence coordinates are designated, if necessary, and sequence algorithm program parameters are designated. Preferably, default program parameters can be used, or alternative parameters can be designated. The sequence comparison algorithm then calculates the percent sequence identities for the test sequences relative to the reference sequence, based on the program parameters,

A "comparison window", as used herein, includes reference to a segment of one of the number of contiguous positions selected from the group consisting typically of from 20 to 600, usually about 50 to about 200, more usually about 100 to about 150 in which a sequence may be compared to a reference sequence of the same number of contiguous positions after the two sequences are optimally aligned. Methods of alignment of sequences for eomparison are well-known in the art. Optimal alignment of sequences for comparison can be conducted, e.g., by the local homology algorithm of Smith & Waterman, Adv. Appl. Math. 2:482 (1981), by the homology alignment algorithm of Needleman & Wunsch, J. Mol. Biol. 48:443 (1970), by the search for similarity method of Pearson & Lipman, Proc. Nat'l. Acad. Sci. USA 85:2444 (1988), by computerized implementations of these algorithms (GAP, BESTFIT, FASTA, and TFASTA in the Wisconsin Genetics Software Package, Genetics Computer Group, 575 Science Dr., Madison, WI), or by manual alignment and visual inspection (see, e.g., Current Protocols in Molecular Biology (Ausubel et al., eds. 1995 supplement)).

Preferred examples of algorithms that are suitable for determining percent sequence identity and sequence similarity include the BLAST and BLAST 2.0 algorithms, which are described in Altschul et al., Nuc. Acids Res. 25:3389-3402 (1977) and Altschul et al., J. Mol. Biol. 215:403-410 (1990). BLAST and BLAST 2.0 are used, with the parameters described herein, to determine percent sequence identity for the nucleic acids and proteins of

the invention. Software for performing BLAST analyses is publicly available through the National Center for Biotechnology Information (http://www.ncbi.nlm.nih.gov/). This algorithm involves first identifying high scoring sequence pairs (HSPs) by identifying short words of length W in the query sequence, which either match or satisfy some positive-valued threshold score T when aligned with a word of the same length in a database sequence. T is referred to as the neighborhood word score threshold (Altschul et al., supra). These initial neighborhood word hits act as seeds for initiating searches to find longer HSPs containing them. The word hits are extended in both directions along each sequence for as far as the cumulative alignment score can be increased. Cumulative scores are calculated using, e.g., for nucleotide sequences, the parameters M (reward score for a pair of matching residues; always > 0) and N (penalty score for mismatching residues; always < 0). For amino acid sequences, a scoring matrix is used to calculate the cumulative score. Extension of the word hits in each direction are halted when: the cumulative alignment score falls off by the quantity X from its maximum achieved value; the cumulative score goes to zero or below, due to the accumulation of one or more negative-scoring residue alignments; or the end of either sequence is reached. The BLAST algorithm parameters W, T, and X determine the sensitivity and speed of the alignment. The BLASTN program (for nucleotide sequences) uses as defaults a wordlength (W) of 11, an expectation (E) of 10, M=5, N=-4 and a comparison of both strands. For amino acid sequences, the BLASTP program uses as defaults a wordlength of 3, and expectation (E) of 10, and the BLOSUM62 scoring matrix (see Henikoff & Henikoff, Proc. Natl. Acad. Sci. USA 89:10915 (1989)) alignments (B) of 50, expectation (E) of 10, M=5, N=-4, and a comparison of both strands.

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The BLAST algorithm also performs a statistical analysis of the similarity between two sequences (see, e.g., Karlin & Altschul, Proc. Nat'l. Acad. Sci. USA 90:5873-5787 (1993)). One measure of similarity provided by the BLAST algorithm is the smallest sum probability (P(N)), which provides an indication of the probability by which a match between two nucleotide or amino acid sequences would occur by chance. For example, a nucleic acid is considered similar to a reference sequence if the smallest sum probability in a comparison of the test nucleic acid to the reference nucleic acid is less than about 0.2, more preferably less than about 0.01, and most preferably less than about 0.001. Log values may be large negative numbers, e.g., 5, 10, 20, 30, 40, 40, 70, 90, 110, 150, 170, etc.

An indication that two nucleic acid sequences or polypeptides are substantially identical is that the polypeptide encoded by the first nucleic acid is immunologically cross reactive with the antibodies raised against the polypeptide encoded by the second nucleic acid, as described below. Thus, a polypeptide is typically substantially identical to a second polypeptide, e.g., where the two peptides differ only by conservative substitutions. Another indication that two nucleic acid sequences are substantially identical is that the two molecules or their complements hybridize to each other under stringent conditions, as described below. Yet another indication that two nucleic acid sequences are substantially identical is that the same primers can be used to amplify the sequences.

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A "host cell" is a naturally occurring cell or a transformed cell that contains an expression vector and supports the replication or expression of the expression vector. Host cells may be cultured cells, explants, cells in vivo, and the like. Host cells may be prokaryotic cells such as E. coli, or eukaryotic cells such as yeast, insect, amphibian, or mammalian cells such as CHO, HeLa, and the like (see, e.g., the American Type Culture Collection catalog or web site, www.atcc.org).

The terms "isolated," "purified," or "biologically pure" refer to material that is substantially or essentially free from components that normally accompany it as found in its native state. Purity and homogeneity are typically determined using analytical chemistry techniques such as polyacrylamide gel electrophoresis or high performance liquid chromatography. A protein or nucleic acid that is the predominant species present in a preparation is substantially purified. In particular, an isolated nucleic acid is separated from some open reading frames that naturally flank the gene and encode proteins other than protein encoded by the gene. The term "purified" in some embodiments denotes that a nucleic acid or protein gives rise to essentially one band in an electrophoretic gel. Preferably, it means that the nucleic acid or protein is at least 85% pure, more preferably at least 95% pure, and most preferably at least 99% pure. "Purify" or "purification" in other embodiments means removing at least one contaminant from the composition to be purified. In this sense, purification does not require that the purified compound be homogenous, e.g., 100% pure.

herein to refer to a polymer of amino acid residues. The terms apply to amino acid polymers in which one or more amino acid residue is an artificial chemical mimetic of a corresponding

The terms "polypeptide." "peptide" and "protein" are used interchangeably

naturally occurring amino acid, as well as to naturally occurring amino acid polymers, those containing modified residues, and non-naturally occurring amino acid polymer.

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The term "amino acid" refers to naturally occurring and synthetic amino acids, as well as amino acid analogs and amino acid mimetics that function similarly to the naturally occurring amino acids. Naturally occurring amino acids are those encoded by the genetic code, as well as those amino acids that are later modified, e.g., hydroxyproline, γ-carboxyglutamate, and O-phosphoserine. Amino acid analogs refers to compounds that have the same basic chemical structure as a naturally occurring amino acid, e.g., an α carbon that is bound to a hydrogen, a carboxyl group, an amino group, and an R group, e.g., homoserine, norleucine, methionine sulfoxide, methionine methyl sulfonium. Such analogs may have modified R groups (e.g., norleucine) or modified peptide backbones, but retain the same basic chemical structure as a naturally occurring amino acid. Amino acid mimetics refers to chemical compounds that have a structure that is different from the general chemical structure of an amino acid, but that functions similarly to a naturally occurring amino acid.

Amino acids may be referred to herein by either their commonly known three letter symbols or by the one-letter symbols recommended by the IUPAC-IUB Biochemical Nomenclature Commission. Nucleotides, likewise, may be referred to by their commonly accepted single-letter codes.

"Conservatively modified variants" applies to both amino acid and nucleic acid sequences. With respect to particular nucleic acid sequences, conservatively modified variants refers to those nucleic acids which encode identical or essentially identical amino acid sequences, or where the nucleic acid does not encode an amino acid sequence, to essentially identical or associated, e.g., naturally contiguous, sequences. Because of the degeneracy of the genetic code, a large number of functionally identical nucleic acids encode most proteins. For instance, the codons GCA, GCC, GCG and GCU all encode the amino acid alanine. Thus, at every position where an alanine is specified by a codon, the codon can be altered to another of the corresponding codons described without altering the encoded polypeptide. Such nucleic acid variations are "silent variations," which are one species of conservatively modified variations. Every nucleic acid sequence herein which encodes a polypeptide also describes silent variations of the nucleic acid. One of skill will recognize that in certain contexts each codon in a nucleic acid (except AUG, which is ordinarily the

only codon for methionine, and TGG, which is ordinarily the only codon for tryptophan) can be modified to yield a functionally identical molecule. Accordingly, often silent variations of a nucleic acid which encodes a polypeptide is implicit in a described sequence with respect to the expression product, but not with respect to actual probe sequences.

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As to amino acid sequences, one of skill will recognize that individual substitutions, deletions or additions to a nucleic acid, peptide, polypeptide, or protein sequence which alters, adds or deletes a single amino acid or a small percentage of amino acids in the encoded sequence is a "conservatively modified variant" where the alteration results in the substitution of an amino acid with a chemically similar amino acid. Conservative substitution tables providing functionally similar amino acids are well known in the art. Such conservatively modified variants are in addition to and do not exclude polymorphic variants, interspecies homologs, and alleles of the invention typically conservative substitutions for one another: 1) Alanine (A), Glycine (G); 2) Aspartic acid (D), Glutamic acid (E); 3) Asparagine (N), Glutamine (O); 4) Arginine (R), Lysine (K); 5) Isoleucine (I), Leucine (L), Methionine (M), Valine (V); 6) Phenylalanine (F), Tyrosine (Y), 15 Tryptophan (W); 7) Serine (S), Threonine (T); and 8) Cysteine (C), Methionine (M) (see, e.g., Creighton, Proteins (1984)).

Macromolecular structures such as polypeptide structures can be described in terms of various levels of organization. For a general discussion of this organization, see, e.g., Alberts et al., Molecular Biology of the Cell (3rd ed., 1994) and Cantor & Schimmel, Biophysical Chemistry Part I: The Conformation of Biological Macromolecules (1980). "Primary structure" refers to the amino acid sequence of a particular peptide. "Secondary structure" refers to locally ordered, three dimensional structures within a polypeptide. These structures are commonly known as domains. Domains are portions of a polypeptide that often form a compact unit of the polypeptide and are typically 25 to approximately 500 amino acids long. Typical domains are made up of sections of lesser organization such as stretches of β-sheet and α-helices. "Tertiary structure" refers to the complete three dimensional structure of a polypeptide monomer. "Quaternary structure" refers to the three dimensional structure formed, usually by the noncovalent association of independent tertiary units. Anisotropic terms are also known as energy terms.

"Nucleic acid" or "oligonucleotide" or "polynucleotide" or grammatical equivalents used herein means at least two nucleotides covalently linked together. Oligonucleotides are typically from about 5, 6, 7, 8, 9, 10, 12, 15, 25, 30, 40, 50 or more nucleotides in length, up to about 100 nucleotides in length. Nucleic acids and polynucleotides are a polymers of any length, including longer lengths, e.g., 200, 300, 500, 1000, 2000, 3000, 5000, 7000, 10,000, etc. A nucleic acid of the present invention will generally contain phosphodiester bonds, although in some cases, nucleic acid analogs are included that may have alternate backbones, comprising, e.g., phosphoramidate, phosphorothioate, phosphorodithioate, or O-methylphophoroamidite linkages (see Eckstein, Oligonucleotides and Analogues: A Practical Approach, Oxford University Press): and peptide nucleic acid backbones and linkages. Other analog nucleic acids include those with positive backbones; non-ionic backbones, and non-ribose backbones, including those described in U.S. Patent Nos. 5,235,033 and 5,034,506, and Chapters 6 and 7, ASC Symposium Series 580, Carbohydrate Modifications in Antisense Research, Sanghui & Cook, eds.. Nucleic acids containing one or more carbocyclic sugars are also included within one definition of nucleic acids. Modifications of the ribose-phosphate backbone may be done for a variety of reasons, e.g. to increase the stability and half-life of such molecules in physiological environments or as probes on a biochip. Mixtures of naturally occurring nucleic acids and analogs can be made; alternatively, mixtures of different nucleic acid analogs, and mixtures of naturally occurring nucleic acids and analogs may be made.

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example, phosphoramidate (Beaucage et al., Tetrahedron 49(10):1925 (1993) and references therein; Letsinger, J. Org. Chem. 35:3800 (1970); Sprinzl et al., Eur. J. Biochem. 81:579 (1977); Letsinger et al., Nucl. Acids Res. 14:3487 (1986); Sawai et al, Chem. Lett. 805 (1984), Letsinger et al., J. Am. Chem. Soc. 110:4470 (1988); and Pauwels et al., Chemica Scripta 26:141 91986)), phosphorothioate (Mag et al., Nucleic Acids Res. 19:1437 (1991); and U.S. Patent No. 5,644,048), phosphorodithioate (Briu et al., J. Am. Chem. Soc. 111:2321 (1989), O-methylphophoroamidite linkages (see Eckstein, Oligonucleotides and Analogues: A Practical Approach, Oxford University Press), and peptide nucleic acid backbones and linkages (see Egholm, J. Am. Chem. Soc. 114:1895 (1992); Meier et al., Chem. Int. Ed. Engl. 31:1008 (1992); Nielsen, Nature, 365:566 (1993); Carlsson et al., Nature 380:207 (1996), all

A variety of references disclose such nucleic acid analogs, including, for

of which are incorporated by reference). Other analog nucleic acids include those with positive backbones (Denpcy et al., Proc. Natl. Acad. Sci. USA 92:6097 (1995); non-ionic backbones (U.S. Patent Nos. 5,386,023, 5,637,684, 5,602,240, 5,216,141 and 4,469,863; Kiedrowshi et al., Angew. Chem. Intl. Ed. English 30:423 (1991); Letsinger et al., J. Am. Chem. Soc. 110:4470 (1988); Letsinger et al., Nucleoside & Nucleotide 13:1597 (1994); Chapters 2 and 3, ASC Symposium Series 580, "Carbohydrate Modifications in Antisense Research", Ed. Y.S. Sanghui and P. Dan Cook; Mesmacker et al., Bioorganic & Medicinal Chem. Lett. 4:395 (1994); Jeffs et al., J. Biomolecular NMR 34:17 (1994); Tetrahedron Lett. 37:743 (1996)) and non-ribose backbones, including those described in U.S. Patent Nos. 5,235,033 and 5,034,506, and Chapters 6 and 7, ASC Symposium Series 580, "Carbohydrate Modifications in Antisense Research", Ed. Y.S. Sanghui and P. Dan Cook. Nucleic acids containing one or more carbocyclic sugars are also included within one definition of nucleic acids (see Jenkins et al., Chem. Soc. Rev. (1995) pp 169-176). Several nucleic acid analogs are described in Rawls, C & E News June 2, 1997 page 35. All of these references are hereby expressly incorporated by reference.

Particularly preferred are peptide nucleic acids (PNA) which includes peptide nucleic acid analogs. These backbones are substantially non-ionic under neutral conditions, in contrast to the highly charged phosphodiester backbone of naturally occurring nucleic acids. This results in two advantages. First, the PNA backbone exhibits improved hybridization kinetics. PNAs have larger changes in the melting temperature ( $T_m$ ) for mismatched versus perfectly matched basepairs. DNA and RNA typically exhibit a 2-4°C drop in  $T_m$  for an internal mismatch. With the non-ionic PNA backbone, the drop is closer to 7-9°C. Similarly, due to their non-ionic nature, hybridization of the bases attached to these backbones is relatively insensitive to salt concentration. In addition, PNAs are not degraded by cellular enzymes, and thus can be more stable.

The nucleic acids may be single stranded or double stranded, as specified, or contain portions of both double stranded or single stranded sequence. As will be appreciated by those in the art, the depiction of a single strand also defines the sequence of the complementary strand; thus the sequences described herein also provide the complement of the sequence. The nucleic acid may be DNA, both genomic and cDNA, RNA or a hybrid, where the nucleic acid may contain combinations of deoxyribo- and ribo-nucleotides, and

combinations of bases, including uracil, adenine, thymine, cytosine, guanine, inosine, xanthine hypoxanthine, isocytosine, isoguanine, etc. "Transcript" typically refers to a naturally occurring RNA, e.g., a pre-mRNA, hnRNA, or mRNA. As used herein, the term "nucleoside" includes nucleotides and nucleoside and nucleotide analogs, and modified nucleosides such as amino modified nucleosides. In addition, "nucleoside" includes non-naturally occurring analog structures. Thus, e.g. the individual units of a peptide nucleic acid, each containing a base, are referred to herein as a nucleoside.

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A "label" or a "detectable moiety" is a composition detectable by spectroscopic, photochemical, biochemical, immunochemical, chemical, or other physical means. For example, useful labels include fluorescent dyes, electron-dense reagents, enzymes (e.g., as commonly used in an ELISA), biotin, digoxigenin, or haptens and proteins or other entities which can be made detectable, e.g., by incorporating a radiolabel into the peptide or used to detect antibodies specifically reactive with the peptide. The radioisotope may be, for example, 3H, 14C, 32P, 35S, or 125I. In some cases, particularly using antibodies against the proteins of the invention, the radioisotopes are used as toxic moieties, as described below. The labels may be incorporated into the prostate cancer nucleic acids, proteins and antibodies at any position. Any method known in the art for conjugating the antibody to the label may be employed, including those methods described by Hunter et al., Nature, 144:945 (1962); David et al., Biochemistry, 13:1014 (1974); Pain et al., J. Immunol, Meth., 40:219 (1981); and Nygren, J. Histochem, and Cytochem., 30:407 (1982). The lifetime of radiolabeled peptides or radiolabeled antibody compositions may extended by the addition of substances that stablize the radiolabeled peptide or antibody and protect it from degradation. Any substance or combination of substances that stablize the radiolabeled peptide or antibody may be used including those substances disclosed in US Patent No. 5,961,955.

An "effector" or "effector moiety" or "effector component" is a molecule that is bound (or linked, or conjugated), either covalently, through a linker or a chemical bond, or noncovalently, through ionic, van der Waals, electrostatic, or hydrogen bonds, to an antibody. The "effector" can be a variety of molecules including, e.g., detection moieties including radioactive compounds, fluorescent compounds, an enzyme or substrate, tags such as epitope tags, a toxin; activatable moieties, a chemotherapeutic agent; a lipase; an antibiotic; or a radioisotope emitting "hard" e.g., beta radiation.

A "labeled nucleic acid probe or oligonucleotide" is one that is bound, either covalently, through a linker or a chemical bond, or noncovalently, through ionic, van der Waals, electrostatic, or hydrogen bonds to a label such that the presence of the probe may be detected by detecting the presence of the label bound to the probe. Alternatively, method using high affinity interactions may achieve the same results where one of a pair of binding partners binds to the other, e.g., biotin, streptavidin.

As used herein a "nucleic acid probe or oligonucleotide" is defined as a nucleic acid capable of binding to a target nucleic acid of complementary sequence through one or more types of chemical bonds, usually through complementary base pairing, usually through hydrogen bond formation. As used herein, a probe may include natural (i.e., A, G, C, or T) or modified bases (7-deazaguanosine, inosine, etc.). In addition, the bases in a probe may be joined by a linkage other than a phosphodiester bond, so long as it does not functionally interfere with hybridization. Thus, e.g., probes may be peptide nucleic acids in which the constituent bases are joined by peptide bonds rather than phosphodiester linkages. It will be understood by one of skill in the art that probes may bind target sequences lacking complete complementarity with the probe sequence depending upon the stringency of the hybridization conditions. The probes are preferably directly labeled as with isotopes, chromophores, lumiphores, chromogens, or indirectly labeled such as with biotin to which a streptavidin complex may later bind. By assaying for the presence or absence of the probe, one can detect the presence or absence of the select sequence or subsequence. Diagnosis or prognosis may be based at the genomic level, or at the level of RNA or protein expression.

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The term "recombinant" when used with reference, e.g., to a cell, or nucleic acid, protein, or vector, indicates that the cell, nucleic acid, protein or vector, has been modified by the introduction of a heterologous nucleic acid or protein or the alteration of a native nucleic acid or protein, or that the cell is derived from a cell so modified. Thus, e.g., recombinant cells express genes that are not found within the native (non-recombinant) form of the cell or express genes that are otherwise abnormally expressed, under expressed or not expressed at all. By the term "recombinant nucleic acid" herein is meant nucleic acid, originally formed in vitro, in general, by the manipulation of nucleic acid, e.g., using polymerases and endonucleases, in a form not normally found in nature. In this manner, operably linkage of different sequences is achieved. Thus an isolated nucleic acid, in a linear

form, or an expression vector formed in vitro by ligating DNA molecules that are not normally joined, are both considered recombinant for the purposes of this invention. It is understood that once a recombinant nucleic acid is made and reintroduced into a host cell or organism, it will replicate non-recombinantly, i.e., using the in vivo cellular machinery of the host cell rather than in vitro manipulations; however, such nucleic acids, once produced recombinantly, although subsequently replicated non-recombinantly, are still considered recombinant for the purposes of the invention. Similarly, a "recombinant protein" is a protein made using recombinant techniques, i.e., through the expression of a recombinant nucleic acid as depicted above.

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The term "heterologous" when used with reference to portions of a nucleic acid indicates that the nucleic acid comprises two or more subsequences that are not normally found in the same relationship to each other in nature. For instance, the nucleic acid is typically recombinantly produced, having two or more sequences, e.g., from unrelated genes arranged to make a new functional nucleic acid, e.g., a promoter from one source and a 15 coding region from another source. Similarly, a heterologous protein will often refer to two or more subsequences that are not found in the same relationship to each other in nature (e.g., a fusion protein).

A "promoter" is defined as an array of nucleic acid control sequences that direct transcription of a nucleic acid. As used herein, a promoter includes necessary nucleic acid sequences near the start site of transcription, such as, in the case of a polymerase II type promoter, a TATA element. A promoter also optionally includes distal enhancer or repressor elements, which can be located as much as several thousand base pairs from the start site of transcription. A "constitutive" promoter is a promoter that is active under most environmental and developmental conditions. An "inducible" promoter is a promoter that is active under environmental or developmental regulation. The term "operably linked" refers to a functional linkage between a nucleic acid expression control sequence (such as a promoter, or array of transcription factor binding sites) and a second nucleic acid sequence, wherein the expression control sequence directs transcription of the nucleic acid corresponding to the second sequence.

An "expression vector" is a nucleic acid construct, generated recombinantly or synthetically, with a series of specified nucleic acid elements that permit transcription of a

particular nucleic acid in a host cell. The expression vector can be part of a plasmid, virus, or nucleic acid fragment. Typically, the expression vector includes a nucleic acid to be transcribed operably linked to a promoter.

The phrase "selectively (or specifically) hybridizes to" refers to the binding,

duplexing, or hybridizing of a molecule only to a particular nucleotide sequence that is

determinative of the presence of the nucleotide sequence, in a heterogeneous population of

nucleic acids and other biologics (e.g., total cellular or library DNA or RNA). Similarly, the

phrase "specifically (or selectively) binds" to an antibody or "specifically (or selectively)

immunoreactive with," when referring to a protein or peptide, refers to a binding reaction that

is determinative of the presence of the protein, in a heterogeneous population of proteins and

other biologics. Thus, under designated immunoassay or nucleic acid hybridization

conditions, the specified antibodies or nucleic acid probes bind to a particular protein

nucleotide sequences at least two times the background and more typically more than 10 to

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Specific binding to an antibody under such conditions requires an antibody that is selected for its specificity for a particular protein. For example, polyclonal antibodies raised to a particular protein, polymorphic variants, alleles, orthologs, and conservatively modified variants, or splice variants, or portions thereof, can be selected to obtain only those polyclonal antibodies that are specifically immunoreactive with the desired prostact cancer protein and not with other proteins. This selection may be achieved by subtracting out antibodies that cross-react with other molecules. A variety of immunoassay formats may be used to select antibodies specifically immunoreactive with a particular protein. For example, solid-phase ELISA immunoassays are routinely used to select antibodies specifically immunoreactive with a protein (see, e.g., Harlow & Lane, Antibodies, A Laboratory Manual (1988) for a description of immunoassay formats and conditions that can be used to determine specific immunoreactivity).

The phrase "stringent hybridization conditions" refers to conditions under which a probe will hybridize to its target subsequence, typically in a complex mixture of nucleic acids, but to no other sequences. Stringent conditions are sequence-dependent and will be different in different circumstances. Longer sequences hybridize specifically at higher temperatures. An extensive guide to the hybridization of nucleic acids is found in

Tijssen, Techniques in Biochemistry and Molecular Biology--Hybridization with Nucleic Probes, "Overview of principles of hybridization and the strategy of nucleic acid assays" (1993). Generally, stringent conditions are selected to be about 5-10°C lower than the thermal melting point (Tm) for the specific sequence at a defined ionic strength pH. The Tm is the temperature (under defined ionic strength, pH, and nucleic concentration) at which 50% 5 of the probes complementary to the target hybridize to the target sequence at equilibrium (as the target sequences are present in excess, at Tm, 50% of the probes are occupied at equilibrium). Stringent conditions will be those in which the salt concentration is less than about 1.0 M sodium ion, typically about 0.01 to 1.0 M sodium ion concentration (or other salts) at pH 7.0 to 8.3 and the temperature is at least about 30°C for short probes (e.g., 10 to 10 50 nucleotides) and at least about 60°C for long probes (e.g., greater than 50 nucleotides). Stringent conditions may also be achieved with the addition of destabilizing agents such as formamide. For selective or specific hybridization, a positive signal is at least two times background, preferably 10 times background hybridization. Exemplary stringent hybridization conditions can be as following: 50% formamide, 5x SSC, and 1% SDS, incubating at 42°C, or, 5x SSC, 1% SDS, incubating at 65°C, with wash in 0.2x SSC, and 0.1% SDS at 65°C. For PCR, a temperature of about 36°C is typical for low stringency amplification, although annealing temperatures may vary between about 32°C and 48°C depending on primer length. For high stringency PCR amplification, a temperature of about 62°C is typical, although high stringency annealing temperatures can range from about 50°C 20 to about 65°C, depending on the primer length and specificity. Typical cycle conditions for both high and low stringency amplifications include a denaturation phase of 90°C - 95°C for 30 sec - 2 min., an annealing phase lasting 30 sec. - 2 min., and an extension phase of about 72°C for 1 - 2 min. Protocols and guidelines for low and high stringency amplification reactions are provided, e.g., in Innis et al. (1990) PCR Protocols, A Guide to Methods and 25

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Nucleic acids that do not hybridize to each other under stringent conditions are still substantially identical if the polypeptides which they encode are substantially identical. This occurs, e.g., when a copy of a nucleic acid is created using the maximum codon degeneracy permitted by the genetic code. In such cases, the nucleic acids typically hybridize

Applications, Academic Press, Inc. N.Y.).

under moderately stringent hybridization conditions. Exemplary "moderately stringent hybridization conditions" include a hybridization in a buffer of 40% formamide, 1 M NaCl, 1% SDS at 37°C, and a wash in 1X SSC at 45°C. A positive hybridization is at least twice background. Those of ordinary skill will readily recognize that alternative hybridization and wash conditions can be utilized to provide conditions of similar stringency. Additional guidelines for determining hybridization parameters are provided in numerous reference, e.g., and Current Protocols in Molecular Biology, ed. Ausubel, et al.

The phrase "functional effects" in the context of assays for testing compounds that modulate activity of a prostate cancer protein includes the determination of a parameter that is indirectly or directly under the influence of the prostate cancer protein or nucleic acid, e.g., a functional, physical, or chemical effect, such as the ability to decrease prostate cancer. It includes ligand binding activity; cell growth on soft agar; anchorage dependence; contact inhibition and density limitation of growth; cellular proliferation; cellular transformation; growth factor or serum dependence; tumor specific marker levels; invasiveness into Matrigel; tumor growth and metastasis in vivo; mRNA and protein expression in cells undergoing metastasis, and other characteristics of prostate cancer cells. "Functional effects" include in vitro, in vivo, and ex vivo activities.

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By "determining the functional effect" is meant assaying for a compound that increases or decreases a parameter that is indirectly or directly under the influence of a prostate cancer protein sequence, e.g., functional, enzymatic, physical and chemical effects. Such functional effects can be measured by any means known to those skilled in the art, e.g., changes in spectroscopic characteristics (e.g., fluorescence, absorbance, refractive index), hydrodynamic (e.g., shape), chromatographic, or solubility properties for the protein, measuring induing activity or binding assays, e.g. binding to antibodies or other ligands, and measuring cellular proliferation. Determination of the functional effect of a compound on prostate cancer can also be performed using prostate cancer assays known to those of skill in the art such as an *in vitro* assays, e.g., cell growth on soft agar; anchorage dependence; contact inhibition and density limitation of growth; cellular proliferation; cellular transformation; growth factor or serum dependence; tumor specific marker levels; invasiveness into Matrigel; tumor growth and metastasis *in vivo*; mRNA and protein

expression in cells undergoing metastasis, and other characteristics of prostate cancer cells. The functional effects can be evaluated by many means known to those skilled in the art, e.g., microscopy for quantitative or qualitative measures of alterations in morphological features, measurement of changes in RNA or protein levels for prostate cancer-associated sequences, measurement of RNA stability, identification of downstream or reporter gene expression (CAT, luciferase,  $\beta$ -gal, GFP and the like), e.g., via chemiluminescence, fluorescence, colorimetric reactions, antibody binding, inducible markers, and ligand binding assays.

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"Inhibitors", "activators", and "modulators" of prostate cancer polynucleotide and polypeptide sequences are used to refer to activating, inhibitory, or modulating molecules or compounds identified using in vitro and in vivo assays of prostate cancer polynucleotide and polypeptide sequences. Inhibitors are compounds that, e.g., bind to, partially or totally block activity, decrease, prevent, delay activation, inactivate, desensitize, or down regulate the activity or expression of prostate cancer proteins, e.g., antagonists. Antisense nucleic acids may seem to inhibit expression and subsequent function of the protein. "Activators" are compounds that increase, open, activate, facilitate, enhance activation, sensitize, agonize, or up regulate prostate cancer protein activity. Inhibitors, activators, or modulators also include genetically modified versions of prostate cancer proteins, e.g., versions with altered activity, as well as naturally occurring and synthetic ligands, antagonists, agonists, antibodies, small chemical molecules and the like. Such assays for inhibitors and activators include, e.g., expressing the prostate cancer protein in vitro, in cells, or cell membranes, applying putative modulator compounds, and then determining the functional effects on activity, as described above. Activators and inhibitors of prostate cancer can also be identified by incubating prostate cancer cells with the test compound and determining increases or decreases in the expression of 1 or more prostate cancer proteins, e.g., 1, 2, 3, 4, 5, 10, 15, 20, 25, 30, 40, 50 or more prostate cancer proteins, such as prostate cancer proteins encoded by the sequences set out in Tables 1-16.

Samples or assays comprising prostate cancer proteins that are treated with a potential activator, inhibitor, or modulator are compared to control samples without the inhibitor, activator, or modulator to examine the extent of inhibition. Control samples (untreated with inhibitors) are assigned a relative protein activity value of 100%. Inhibition of a polymentide is achieved when the activity value relative to the control is about 80%.

preferably 50%, more preferably 25-0%. Activation of a prostate cancer polypeptide is achieved when the activity value relative to the control (untreated with activators) is 110%, more preferably 150%, more preferably 200-500% (i.e., two to five fold higher relative to the control), more preferably 1000-3000% higher.

The phrase "changes in cell growth" refers to any change in cell growth and proliferation characteristics in vitro or in vivo, such as formation of foci, anchorage independence, semi-solid or soft agar growth, changes in contact inhibition and density limitation of growth, loss of growth factor or scrum requirements, changes in cell morphology, gaining or losing immortalization, gaining or losing tumor specific markers, ability to form or suppress tumors when injected into suitable animal hosts, and/or immortalization of the cell. See, e.g., Freshney, Culture of Animal Cells a Manual of Basic Technique pp. 231-241 (3<sup>rd</sup> ed. 1994).

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"Cancer cells," "transformed" cells or "transformation" in tissue culture, refers to spontaneous or induced phenotypic changes that do not necessarily involve the uptake of new genetic material. Although transformation can arise from infection with a transforming virus and incorporation of new genomic DNA, or uptake of exogenous DNA, it can also arise spontaneously or following exposure to a carcinogen, thereby mutating an endogenous gene. Transformation is associated with phenotypic changes, such as immortalization of cells, aberrant growth control, nonmorphological changes, and/or malignancy (see, Freshney,

"Tumor cell" refers to precancerous, cancerous, and normal cells in a tumor.

"Antibody" refers to a polypeptide comprising a framework region from an immunoglobulin gene or fragments thereof that specifically binds and recognizes an antigen. The recognized immunoglobulin genes include the kappa, lambda, alpha, gamma, delta, epsilon, and mu constant region genes, as well as the myriad immunoglobulin variable region genes. Light chains are classified as either kappa or lambda. Heavy chains are classified as gamma, mu, alpha, delta, or epsilon, which in turn define the immunoglobulin classes, IgG, IgM, IgA, IgD and IgE, respectively. Typically, the antigen-binding region of an antibody or its functional equivalent will be most critical in specificity and affinity of binding. See Paul, Fundamental Immunology.

Culture of Animal Cells a Manual of Basic Technique (3rd ed. 1994)).

An exemplary immunoglobulin (antibody) structural unit comprises a tetramer. Each tetramer is composed of two identical pairs of polypeptide chains, each pair having one "light" (about 25 kD) and one "heavy" chain (about 50-70 kD). The N-terminus of each chain defines a variable region of about 100 to 110 or more amino acids primarily responsible for antigen recognition. The terms variable light chain (V<sub>1</sub>) and variable heavy chain (V<sub>H</sub>) refer to these light and heavy chains respectively.

Antibodies exist, e.g., as intact immunoglobulins or as a number of well-characterized fragments produced by digestion with various peptidases. Thus, e.g., pepsin digests an antibody below the disulfide linkages in the hinge region to produce F(ab)'<sub>2</sub>, a dimer of Fab which itself is a light chain joined to V<sub>H</sub>-C<sub>H</sub>1 by a disulfide bond. The F(ab)'<sub>2</sub> may be reduced under mild conditions to break the disulfide linkage in the hinge region, thereby converting the F(ab)'<sub>2</sub> dimer into an Fab' monomer. The Fab' monomer is essentially Fab with part of the hinge region (see Fundamental Immunology (Paul ed., 3d ed. 1993). While various antibody fragments are defined in terms of the digestion of an intact antibody, one of skill will appreciate that such fragments may be synthesized de novo either chemically or by using recombinant DNA methodology. Thus, the term antibody, as used herein, also includes antibody fragments either produced by the modification of whole antibodies, or those synthesized de novo using recombinant DNA methodologies (e.g., single chain Fv) or those identified using phage display libraries (see, e.g., McCafferty et al., Nature 348:552-554 (1990))

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For preparation of antibodies, e.g., recombinant, monoclonal, or polyclonal antibodies, many technique known in the art can be used (see, e.g., Kohler & Milstein, Nature 256:495-497 (1975); Kozbor et al., Immunology Today 4:72 (1983); Cole et al., pp. 77-96 in Monoclonal Antibodies and Cancer Therapy (1985); Coligan, Current Protocols in Immunology (1991); Harlow & Lane, Antibodies, A Laboratory Manual (1988); and Goding, Monoclonal Antibodies: Principles and Practice (2d ed. 1986)). Techniques for the production of single chain antibodies (U.S. Patent 4,946,778) can be adapted to produce antibodies to polypeptides of this invention. Also, transgenic mice, or other organisms such as other mammals, may be used to express humanized antibodies. Alternatively, phage display technology can be used to identify antibodies and heteromeric Fab fragments that

specifically bind to selected antigens (see, e.g., McCafferty et al., Nature 348:552-554 (1990); Marks et al., Biotechnology 10:779-783 (1992)).

A "chimeric antibody" is an antibody molecule in which (a) the constant region, or a portion thereof, is altered, replaced or exchanged so that the antigen binding site (variable region) is linked to a constant region of a different or altered class, effector function and/or species, or an entirely different molecule which confers new properties to the chimeric antibody, e.g., an enzyme, toxin, hormone, growth factor, drug, etc.; or (b) the variable region, or a portion thereof, is altered, replaced or exchanged with a variable region having a different or altered antigen specificity.

## Identification of prostate cancer-associated sequences

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In one aspect, the expression levels of genes are determined in different patient samples for which diagnosis information is desired, to provide expression profiles. An expression profile of a particular sample is essentially a "fingerprint" of the state of the sample; while two states may have any particular gene similarly expressed, the evaluation of a number of genes simultaneously allows the generation of a gene expression profile that is characteristic of the state of the cell. That is, normal tissue (e.g., normal prostate or other tissue) may be distinguished from cancerous or metastatic cancerous tissue of the prostate, or prostate cancer tissue or metastatic prostate cancerous tissue of the prostate samples of prostate and other tissues from surviving cancer patients. By comparing expression profiles of tissue in known different prostate cancer states, information regarding which genes are important (including both up- and down-regulation of genes) in each of these states is obtained.

The identification of sequences that are differentially expressed in prostate cancer versus non-prostate cancer tissue allows the use of this information in a number of ways. For example, a particular treatment regime may be evaluated: does a chemotherapeutic drug act to down-regulate prostate eancer, and thus tumor growth or recurrence, in a particular patient. Similarly, diagnosis and treatment outcomes may be done or confirmed by comparing patient samples with the known expression profiles. Metastatic tissue can also be analyzed to determine the stage of prostate cancer in the tissue. Furthermore, these gene expression profiles (or individual genes) allow screening of drug candidates with an eye to

mimicking or altering a particular expression profile; e.g., screening can be done for drugs that suppress the prostate cancer expression profile. This may be done by making biochips comprising sets of the important prostate cancer genes, which can then be used in these screens. These methods can also be done on the protein basis; that is, protein expression levels of the prostate cancer proteins can be evaluated for diagnostic purposes or to screen candidate agents. In addition, the prostate cancer nucleic acid sequences can be administered for gene therapy purposes, including the administration of antisense nucleic acids, or the prostate cancer proteins (including antibodies and other modulators thereof) administered as therapeutic drugs.

Thus the present invention provides nucleic acid and protein sequences that are differentially expressed in prostate cancer, herein termed "prostate cancer sequences." As outlined below, prostate cancer sequences include those that are up-regulated (i.e., expressed at a higher level) in prostate cancer, as well as those that are down-regulated (i.e., expressed at a lower level). In a preferred embodiment, the prostate cancer sequences are from humans; however, as will be appreciated by those in the art, prostate cancer sequences from other organisms may be useful in animal models of disease and drug evaluation; thus, other prostate cancer sequences are provided, from vertebrates, including mammals, including rodents (rats, mice, hamsters, guinea pigs, etc.), primates, farm animals (including sheep, goats, pigs, cows, horses, etc.) and pets, e.g., (dogs, cats, etc.). Prostate cancer sequences from other organisms may be obtained using the techniques outlined below.

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Prostate cancer sequences can include both nucleic acid and amino acid sequences. As will be appreciated by those in the art and is more fully outlined below, prostate cancer nucleic acid sequences are useful in a variety of applications, including diagnostic applications, which will detect naturally occurring nucleic acids, as well as screening applications; e.g., biochips comprising nucleic acid probes or PCR microtiter plates with selected probes to the prostate cancer sequences can be generated.

A prostate cancer sequence can be initially identified by substantial nucleic acid and/or amino acid sequence homology to the prostate cancer sequences outlined herein. Such homology can be based upon the overall nucleic acid or amino acid sequence, and is generally determined as outlined below, using either homology programs or hybridization conditions.

For identifying prostate cancer-associated sequences, the prostate cancer screen typically includes comparing genes identified in different tissues, e.g., normal and cancerous tissues, or tumor tissue samples from patients who have metastatic disease vs. non metastatic tissue. Other suitable tissue comparisons include comparing prostate cancer samples with metastatic cancer samples from other cancers, such as lung, breast, gastrointestinal cancers, ovarian, etc. Samples of different stages of prostate cancer, e.g., survivor tissue, drug resistant states, and tissue undergoing metastasis, are applied to biochips comprising nucleic acid probes. The samples are first microdissected, if applicable, and treated as is known in the art for the preparation of mRNA. Suitable biochips are commercially available, e.g. from Affymetrix. Gene expression profiles as described herein are generated and the data analyzed.

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In one embodiment, the genes showing changes in expression as between normal and disease states are compared to genes expressed in other normal tissues, preferably normal prostate, but also including, and not limited to lung, heart, brain, liver, breast, kidney, muscle, colon, small intestine, large intestine, spleen, bone and placenta. In a preferred embodiment, those genes identified during the prostate cancer screen that are expressed in any significant amount in other tissues are removed from the profile, although in some embodiments, this is not necessary. That is, when screening for drugs, it is usually preferable that the target be disease specific, to minimize possible side effects.

In a preferred embodiment, prostate cancer sequences are those that are upregulated in prostate cancer; that is, the expression of these genes is higher in the prostate
cancer tissue as compared to non-cancerous tissue. "Up-regulation" as used herein often
means at least about a two-fold change, preferably at least about a three fold change, with at
least about five-fold or higher being preferred. All unigene cluster identification numbers
and accession numbers herein are for the GenBank sequence database and the sequences of
the accession numbers are hereby expressly incorporated by reference. GenBank is known in
the art, see, e.g., Benson, DA, et al., Nucleic Acids Research 26:1-7 (1998) and
http://www.ncbi.nlm.nih.gov/. Sequences are also available in other databases, e.g.,
European Molecular Biology Laboratory (EMBL) and DNA Database of Japan (DDBJ).

In another preferred embodiment, prostate cancer sequences are those that are down-regulated in prostate cancer; that is, the expression of these genes is lower in prostate

cancer tissue as compared to non-cancerous tissue (see, e.g., Tables 8, 12 and 14). "Down-regulation" as used herein often means at least about a 1.5-fold change more preferrably a two-fold change, preferably at least about a three fold change, with at least about five-fold or higher being most preferred.

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#### Informatics

The ability to identify genes that are over or under expressed in prostate cancer can additionally provide high-resolution, high-sensitivity datasets which can be used in the areas of diagnostics, therapeutics, drug development, pharmacogenetics, protein structure, biosensor development, and other related areas. For example, the expression profiles can be used in diagnostic or prognostic evaluation of patients with prostate cancer. Or as another example, subcellular toxicological information can be generated to better direct drug structure and activity correlation (see Anderson, Pharmaceutical Proteomics: Targets, Mechanism, and Function, paper presented at the IBC Proteomics conference, Coronado, CA (June 11-12, 1998)). Subcellular toxicological information can also be utilized in a biological sensor device to predict the likely toxicological effect of chemical exposures and likely tolerable exposure thresholds (see U.S. Patent No. 5,811,231). Similar advantages accrue from datasets relevant to other biomolecules and bioactive agents (e.g., nucleic acids, saccharides, lipids, drugs, and the like).

Thus, in another embodiment, the present invention provides a database that includes at least one set of assay data. The data contained in the database is acquired, e.g., using array analysis either singly or in a library format. The database can be in substantially any form in which data can be maintained and transmitted, but is preferably an electronic database. The electronic database of the invention can be maintained on any electronic device allowing for the storage of and access to the database, such as a personal computer, but is preferably distributed on a wide area network, such as the World Wide Web.

The focus of the present section on databases that include peptide sequence data is for clarity of illustration only. It will be apparent to those of skill in the art that similar databases can be assembled for any assay data acquired using an assay of the invention.

The compositions and methods for identifying and/or quantitating the relative and/or absolute abundance of a variety of molecular and macromolecular species from a biological sample undergoing prostate cancer, i.e., the identification of prostate cancerassociated sequences described herein, provide an abundance of information, which can be correlated with pathological conditions, predisposition to disease, drug testing, therapeutic monitoring, gene-disease causal linkages, identification of correlates of immunity and physiological status, among others. Although the data generated from the assays of the invention is suited for manual review and analysis, in a preferred embodiment, prior data processing using high-speed computers is utilized.

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An array of methods for indexing and retrieving biomolecular information is known in the art. For example, U.S. Patents 6,023,659 and 5,966,712 disclose a relational database system for storing biomolecular sequence information in a manner that allows sequences to be catalogued and searched according to one or more protein function hierarchies. U.S. Patent 5,953,727 discloses a relational database having sequence records containing information in a format that allows a collection of partial-length DNA sequences 15 to be catalogued and searched according to association with one or more sequencing projects for obtaining full-length sequences from the collection of partial length sequences. U.S. Patent 5,706,498 discloses a gene database retrieval system for making a retrieval of a gene sequence similar to a sequence data item in a gene database based on the degree of similarity between a key sequence and a target sequence. U.S. Patent 5,538,897 discloses a method using mass spectroscopy fragmentation patterns of peptides to identify amino acid sequences in computer databases by comparison of predicted mass spectra with experimentally-derived mass spectra using a closeness-of-fit measure. U.S. Patent 5,926,818 discloses a multidimensional database comprising a functionality for multi-dimensional data analysis described as on-line analytical processing (OLAP), which entails the consolidation of 25 projected and actual data according to more than one consolidation path or dimension. U.S. Patent 5,295,261 reports a hybrid database structure in which the fields of each database record are divided into two classes, navigational and informational data, with navigational fields stored in a hierarchical topological map which can be viewed as a tree structure or as the merger of two or more such tree structures.

See also Mount et al., Bioinformatics (2001); Biological Sequence Analysis: Probabilistic Models of Proteins and Nucleic Acids (Durbin et al., eds., 1999); Bioinformatics: A Practical Guide to the Analysis of Genes and Proteins (Baxevanis & Oeullette eds., 1998)); Rashidi & Buehler, Bioinformatics: Basic Applications in Biological Science and Medicine (1999); Introduction to Computational Molecular Biology (Setubal et al., eds 1997); Bioinformatics: Methods and Protocols (Misener & Krawetz, eds, 2000); Bioinformatics: Sequence, Structure, and Databanks: A Practical Approach (Higgins & Taylor, eds., 2000); Brown, Bioinformatics: A Biologist's Guide to Biocomputing and the Internet (2001); Han & Kamber, Data Mining: Concepts and Techniques (2000); and Waterman, Introduction to Computational Biology: Maps, Sequences, and Genomes (1995).

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The present invention provides a computer database comprising a computer and software for storing in computer-retrievable form assay data records cross-tabulated, e.g., with data specifying the source of the target-containing sample from which each sequence specificity record was obtained.

In an exemplary embodiment, at least one of the sources of target-containing sample is from a control tissue sample known to be free of pathological disorders. In a variation, at least one of the sources is a known pathological tissue specimen, e.g., a neoplastic lesion or another tissue specimen to be analyzed for prostate cancer. In another variation, the assay records cross-tabulate one or more of the following parameters for each target species in a sample: (1) a unique identification code, which can include, e.g., a target molecular structure and/or characteristic separation coordinate (e.g., electrophoretic coordinates); (2) sample source; and (3) absolute and/or relative quantity of the target species present in the sample.

The invention also provides for the storage and retrieval of a collection of target data in a computer data storage apparatus, which can include magnetic disks, optical disks, magneto-optical disks, DRAM, SRAM, SGRAM, SDRAM, RDRAM, DDR RAM, magnetic bubble memory devices, and other data storage devices, including CPU registers and on-CPU data storage arrays. Typically, the target data records are stored as a bit pattern in an array of magnetic domains on a magnetizable medium or as an array of charge states or transistor gate states, such as an array of cells in a DRAM device (e.g., each cell comprised of a transistor and a charge storage area, which may be on the transistor). In one embodiment,

the invention provides such storage devices, and computer systems built therewith, comprising a bit pattern encoding a protein expression fingerprint record comprising unique identifiers for at least 10 target data records cross-tabulated with target source.

When the target is a peptide or nucleic acid, the invention preferably provides

a method for identifying related peptide or nucleic acid sequences, comprising performing a

computerized comparison between a peptide or nucleic acid sequence assay record stored in

or retrieved from a computer storage device or database and at least one other sequence. The

comparison can include a sequence analysis or comparison algorithm or computer program

embodiment thereof (e.g., FASTA, TFASTA, GAP, BESTFIT) and/or the comparison may

be of the relative amount of a peptide or nucleic acid sequence in a pool of sequences

determined from a polypeptide or nucleic acid sample of a specimen.

The invention also preferably provides a magnetic disk, such as an IBM-compatible (DOS, Windows, Windows)5/98/2000, Windows NT, OS/2) or other format (e.g., Linux, SunOS, Solaris, AIX, SCO Unix, VMS, MV, Macintosh, etc.) floppy diskette or hard (fixed, Winchester) disk drive, comprising a bit pattern encoding data from an assay of the invention in a file format suitable for retrieval and processing in a computerized sequence analysis, comparison, or relative quantitation method.

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The invention also provides a network, comprising a plurality of computing devices linked via a data link, such as an Ethernet cable (coax or 10BaseT), telephone line, ISDN line, wireless network, optical fiber, or other suitable signal transmission medium, whereby at least one network device (e.g., computer, disk array, etc.) comprises a pattern of magnetic domains (e.g., magnetic disk) and/or charge domains (e.g., an array of DRAM cells) composing a bit pattern encoding data acquired from an assay of the invention.

The invention also provides a method for transmitting assay data that includes generating an electronic signal on an electronic communications device, such as a modem, ISDN terminal adapter, DSL, cable modem, ATM switch, or the like, wherein the signal includes (in native or encrypted format) a bit pattern encoding data from an assay or a database comprising a plurality of assay results obtained by the method of the invention.

In a preferred embodiment, the invention provides a computer system for comparing a query target to a database containing an array of data structures, such as an assay result obtained by the method of the invention, and ranking database targets based on the

degree of identity and gap weight to the target data. A central processor is preferably initialized to load and execute the computer program for alignment and/or comparison of the assay results. Data for a query target is entered into the central processor via an I/O device. Execution of the computer program results in the central processor retrieving the assay data from the data file, which comprises a binary description of an assay result.

The target data or record and the computer program can be transferred to secondary memory, which is typically random access memory (e.g., DRAM, SRAM, SGRAM, or SDRAM). Targets are ranked according to the degree of correspondence between a selected assay characteristic (e.g., binding to a selected affinity moiety) and the same characteristic of the query target and results are output via an I/O device. For example, a central processor can be a conventional computer (e.g., Intel Pentium, PowerPC, Alpha, PA-8000, SPARC, MIPS 4400, MIPS 10000, VAX, etc.); a program can be a commercial or public domain molecular biology software package (e.g., UWGCG Sequence Analysis Software, Darwin); a data file can be an optical or magnetic disk, a data server, a memory device (e.g., DRAM, SRAM, SGRAM, SDRAM, EPROM, bubble memory, flash memory, etc.); an I/O device can be a terminal comprising a video display and a keyboard, a modem, an ISDN terminal adapter, an Ethernet port, a punched card reader, a magnetic strip reader, or other suitable I/O device.

The invention also preferably provides the use of a computer system, such as that described above, which comprises: (1) a computer; (2) a stored bit pattern encoding a collection of peptide sequence specificity records obtained by the methods of the invention, which may be stored in the computer; (3) a comparison target, such as a query target; and (4) a program for alignment and comparison, typically with rank-ordering of comparison results on the basis of computed similarity values.

# Characteristics of prostate cancer-associated proteins

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Prostate cancer proteins of the present invention may be classified as secreted proteins, transmembrane proteins or intracellular proteins. In one embodiment, the prostate cancer protein is an intracellular protein. Intracellular proteins may be found in the cytoplasm and/or in the nucleus. Intracellular proteins are involved in all aspects of cellular function and replication (including, e.g., signaling pathways); aberrant expression of such

proteins often results in unregulated or disregulated cellular processes (see, e.g., Molecular Biology of the Cell (Alberts, ed., 3rd ed., 1994). For example, many intracellular proteins have enzymatic activity such as protein kinase activity, protein phosphatase activity, protease activity, nucleotide cyclase activity, polymerase activity and the like. Intracellular proteins also serve as docking proteins that are involved in organizing complexes of proteins, or targeting proteins to various subcellular localizations, and are involved in maintaining the structural integrity of organelles.

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(1998)).

An increasingly appreciated concept in characterizing proteins is the presence in the proteins of one or more motifs for which defined functions have been attributed. In addition to the highly conserved sequences found in the enzymatic domain of proteins, highly conserved sequences have been identified in proteins that are involved in protein-protein interaction. For example, Src-homology-2 (SH2) domains bind tyrosine-phosphorylated targets in a sequence dependent manner. PTB domains, which are distinct from SH2 domains, also bind tyrosine phosphorylated targets. SH3 domains bind to proline-rich targets. In addition, PH domains, tetratricopeptide repeats and WD domains to name only a few, have been shown to mediate protein-protein interactions. Some of these may also be involved in binding to phospholipids or other second messengers. As will be appreciated by one of ordinary skill in the art, these motifs can be identified on the basis of primary sequence; thus, an analysis of the sequence of proteins may provide insight into both the enzymatic potential of the molecule and/or molecules with which the protein may associate. One useful database is Pfam (protein families), which is a large collection of multiple sequence alignments and hidden Markov models covering many common protein domains. Versions are available via the internet from Washington University in St. Louis, the Sanger Center in England, and the Karolinska Institute in Sweden (see, e.g., Bateman et al., Nuc. Acids Res. 28:263-266 (2000); Sonnhammer et al., Proteins 28:405-420 (1997); Bateman et al., Nuc. Acids Res. 27:260-262 (1999); and Sonnhammer et al., Nuc. Acids Res. 26:320-322-

In another embodiment, the prostate cancer sequences are transmembrane proteins. Transmembrane proteins are molecules that span a phospholipid bilayer of a cell. They may have an intracellular domain, an extracellular domain, or both. The intracellular domains of such proteins may have a number of functions including those already described

for intracellular proteins. For example, the intracellular domain may have enzymatic activity and/or may serve as a binding site for additional proteins. Frequently the intracellular domain of transmembrane proteins serves both roles. For example certain receptor tyrosine kinases have both protein kinase activity and SH2 domains. In addition, autophosphorylation of tyrosines on the receptor molecule itself, creates binding sites for additional SH2 domain containing proteins.

Transmembrane proteins may contain from one to many transmembrane domains. For example, receptor tyrosine kinases, certain cytokine receptors, receptor guanvlyl cyclases and receptor serine/threonine protein kinases contain a single transmembrane domain. However, various other proteins including channels and adenylyl cyclases contain numerous transmembrane domains. Many important cell surface receptors such as G protein coupled receptors (GPCRs) are classified as "seven transmembrane domain" proteins, as they contain 7 membrane spanning regions. Characteristics of transmembrane domains include approximately 20 consecutive hydrophobic amino acids that may be followed by charged amino acids. Therefore, upon analysis of the amino acid sequence of a particular protein, the localization and number of transmembrane domains within the protein may be predicted (see, e.g. PSORT web site http://psort.nibb.ac.jp/). Important transmembrane protein receptors include, but are not limited to the insulin receptor, insulin-like growth factor receptor, human growth hormone receptor, glucose transporters, transferrin receptor, epidermal growth factor receptor, low density lipoprotein receptor, epidermal growth factor receptor, leptin receptor, interleukin receptors, e.g. IL-1 receptor, IL-2 receptor,

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The extracellular domains of transmembrane proteins are diverse; however, conserved motifs are found repeatedly among various extracellular domains. Conserved structure and/or functions have been ascribed to different extracellular motifs. Many extracellular domains are involved in binding to other molecules. In one aspect, extracellular domains are found on receptors. Factors that bind the receptor domain include circulating ligands, which may be peptides, proteins, or small molecules such as adenosine and the like. For example, growth factors such as EGF, FGF and PDGF are circulating growth factors that bind to their cognate receptors to initiate a variety of cellular responses. Other factors include cytokines, mitogenic factors, neurotrophic factors and the like. Extracellular domains also

bind to cell-associated molecules. In this respect, they mediate cell-cell interactions, Cell-associated ligands can be tethered to the cell, e.g., via a glycosylphosphatidylinositol (GPI) anchor, or may themselves be transmembrane proteins. Extracellular domains also associate with the extracellular matrix and contribute to the maintenance of the cell structure.

Prostate cancer proteins that are transmembrane are particularly preferred in the present invention as they are readily accessible targets for immunotherapeutics, as are described herein. In addition, as outlined below, transmembrane proteins can be also useful in imaging modalities. Antibodies may be used to label such readily accessible proteins in situ. Alternatively, antibodies can also label intracellular proteins, in which case samples are typically permeablized to provide access to intracellular proteins.

It will also be appreciated by those in the art that a transmembrane protein can be made soluble by removing transmembrane sequences, e.g., through recombinant methods. Furthermore, transmembrane proteins that have been made soluble can be made to be secreted through recombinant means by adding an appropriate signal sequence.

In another embodiment, the prostate cancer proteins are secreted proteins; the secretion of which can be either constitutive or regulated. These proteins have a signal peptide or signal sequence that targets the molecule to the secretory pathway. Secreted proteins are involved in numerous physiological events; by virtue of their circulating nature, they serve to transmit signals to various other cell types. The secreted protein may function in an autocrine manner (acting on the cell that secreted the factor), a paracrine manner (acting on cells in close proximity to the cell that secreted the factor) or an endocrine manner (acting on cells at a distance). Thus secreted molecules find use in modulating or altering numerous aspects of physiology. Prostate cancer proteins that are secreted proteins are particularly preferred in the present invention as they serve as good targets for diagnostic markers, e.g., for blood, plasma, serum, or stool tests.

### Use of prostate cancer nucleic acids

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As described above, prostate cancer sequence is initially identified by substantial nucleic acid and/or amino acid sequence homology or linkage to the prostate cancer sequences outlined herein. Such homology can be based upon the overall nucleic acid or amino acid sequence, and is generally determined as outlined below, using either

homology programs or hybridization conditions. Typically, linked sequences on a mRNA are found on the same molecule.

The prostate cancer nucleic acid sequences of the invention, e.g., the sequences in Tables 1-16, can be fragments of larger genes, i.e., they are nucleic acid segments. "Genes" in this context includes coding regions, non-coding regions, and mixtures of coding and non-coding regions. Accordingly, as will be appreciated by those in the art, using the sequences provided herein, extended sequences, in either direction, of the prostate cancer genes can be obtained, using techniques well known in the art for cloning either longer sequences or the full length sequences; see Ausubel, et al., supra. Much can be done by informatics and many sequences can be clustered to include multiple sequences corresponding to a single gene, e.g., systems such as UniGene (see, http://www.ncbi.nlm.nih.gov/UniGene/).

Once the prostate cancer nucleic acid is identified, it can be cloned and, if necessary, its constituent parts recombined to form the entire prostate cancer nucleic acid coding regions or the entire mRNA sequence. Once isolated from its natural source, e.g., contained within a plasmid or other vector or excised therefrom as a linear nucleic acid segment, the recombinant prostate cancer nucleic acid can be further-used as a probe to identify and isolate other prostate cancer nucleic acids, e.g., extended coding regions. It can also be used as a "precursor" nucleic acid to make modified or variant prostate cancer nucleic acids and proteins.

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The prostate cancer nucleic acids of the present invention are used in several ways. In a first embodiment, nucleic acid probes to the prostate cancer nucleic acids are made and attached to biochips to be used in screening and diagnostic methods, as outlined below, or for administration, e.g., for gene therapy, vaccine, and/or antisense applications. Alternatively, the prostate cancer nucleic acids that include coding regions of prostate cancer proteins can be put into expression vectors for the expression of prostate cancer proteins, again for screening purposes or for administration to a patient.

In a preferred embodiment, nucleic acid probes to prostate cancer nucleic acids (both the nucleic acid sequences outlined in the figures and/or the complements thereof) are made. The nucleic acid probes attached to the biochip are designed to be substantially complementary to the prostate cancer nucleic acids, i.e. the target sequence (either the target

sequence of the sample or to other probe sequences, e.g., in sandwich assays), such that hybridization of the target sequence and the probes of the present invention occurs. As outlined below, this complementarity need not be perfect; there may be any number of base pair mismatches which will interfere with hybridization between the target sequence and the single stranded nucleic acids of the present invention. However, if the number of mutations is so great that no hybridization can occur under even the least stringent of hybridization conditions, the sequence is not a complementary target sequence. Thus, by "substantially complementary" herein is meant that the probes are sufficiently complementary to the target sequences to hybridize under normal reaction conditions, particularly high stringency conditions, as outlined herein.

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A nucleic acid probe is generally single stranded but can be partially single and partially double stranded. The strandedness of the probe is dictated by the structure, composition, and properties of the target sequence. In general, the nucleic acid probes range from about 8 to about 100 bases long, with from about 10 to about 80 bases being preferred, and from about 30 to about 50 bases being particularly preferred. That is, generally whole genes are not used. In some embodiments, much longer nucleic acids can be used, up to hundreds of bases.

In a preferred embodiment, more than one probe per sequence is used, with either overlapping probes or probes to different sections of the target being used. That is, two, three, four or more probes, with three being preferred, are used to build in a redundancy for a particular target. The probes can be overlapping (i.e., have some sequence in common), or separate. In some cases, PCR primers may be used to amplify signal for higher sensitivity.

As will be appreciated by those in the art, nucleic acids can be attached or immobilized to a solid support in a wide variety of ways. By "immobilized" and grammatical equivalents herein is meant the association or binding between the nucleic acid probe and the solid support is sufficient to be stable under the conditions of binding, washing, analysis, and removal as outlined below. The binding can typically be covalent or non-covalent. By "non-covalent binding" and grammatical equivalents herein is meant one or more of electrostatic, hydrophilic, and hydrophobic interactions. Included in non-covalent binding is the covalent attachment of a molecule, such as, streptavidin to the support and the non-covalent binding of the biotinylated probe to the streptavidin. By "covalent binding" and grammatical

equivalents herein is meant that the two moieties, the solid support and the probe, are attached by at least one bond, including sigma bonds, pi bonds and coordination bonds. Covalent bonds can be formed directly between the probe and the solid support or can be formed by a cross linker or by inclusion of a specific reactive group on either the solid support or the probe or both molecules. Immobilization may also involve a combination of covalent and non-covalent interactions.

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In general, the probes are attached to the biochip in a wide variety of ways, as will be appreciated by those in the art. As described herein, the nucleic acids can either be synthesized first, with subsequent attachment to the biochip, or can be directly synthesized on the biochip.

The biochip comprises a suitable solid substrate. By "substrate" or "solid support" or other grammatical equivalents herein is meant a material that can be modified to contain discrete individual sites appropriate for the attachment or association of the nucleic acid probes and is amenable to at least one detection method. As will be appreciated by those in the art, the number of possible substrates are very large, and include, but are not limited to, glass and modified or functionalized glass, plastics (including acrylics, polystyrene and copolymers of styrene and other materials, polypropylene, polyethylene, polybutylene, polyurethanes, TeflonJ, etc.), polysaccharides, nylon or nitrocellulose, resins, silica or silicabased materials including silicon and modified silicon, carbon, metals, inorganic glasses, plastics, etc. In general, the substrates allow optical detection and do not appreciably fluoresce. A preferred substrate is described in copending application entitled Reusable Low Fluorescent Plastic Biochip, U.S. Application Serial No. 09/270,214, filed March 15, 1999, herein incorporated by reference in its entirety.

Generally the substrate is planar, although as will be appreciated by those in the art, other configurations of substrates may be used as well. For example, the probes may be placed on the inside surface of a tube, for flow-through sample analysis to minimize sample volume. Similarly, the substrate may be flexible, such as a flexible foam, including closed cell foams made of particular plastics.

In a preferred embodiment, the surface of the biochip and the probe may be derivatized with chemical functional groups for subsequent attachment of the two. Thus, e.g., the biochip is derivatized with a chemical functional group including, but not limited to,

amino groups, carboxy groups, oxo groups and thiol groups, with amino groups being particularly preferred. Using these functional groups, the probes can be attached using functional groups on the probes. For example, nucleic acids containing amino groups can be attached to surfaces comprising amino groups, e.g. using linkers as are known in the art; e.g., homo-or hetero-bifunctional linkers as are well known (see 1994 Pierce Chemical Company catalog, technical section on cross-linkers, pages 155-200). In addition, in some cases, additional linkers, such as alkyl groups (including substituted and heteroalkyl groups) may be used

In this embodiment, oligonucleotides are synthesized as is known in the art, and then attached to the surface of the solid support. As will be appreciated by those skilled in the art, either the 5' or 3' terminus may be attached to the solid support, or attachment may be via an internal nucleoside.

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In another embodiment, the immobilization to the solid support may be very strong, yet non-covalent. For example, biotinylated oligonucleotides can be made, which bind to surfaces covalently coated with streptavidin, resulting in attachment.

Alternatively, the oligonucleotides may be synthesized on the surface, as is known in the art. For example, photoactivation techniques utilizing photopolymerization compounds and techniques are used. In a preferred embodiment, the nucleic acids can be synthesized in situ, using well known photolithographic techniques, such as those described in WO 95/25116; WO 95/35505; U.S. Patent Nos. 5,700,637 and 5,445,934; and references cited within, all of which are expressly incorporated by reference; these methods of attachment form the basis of the Affimetrix GeneChip<sup>TM</sup> technology.

Often, amplification-based assays are performed to measure the expression level of prostate cancer-associated sequences. These assays are typically performed in conjunction with reverse transcription. In such assays, a prostate cancer-associated nucleic acid sequence acts as a template in an amplification reaction (e.g., Polymerase Chain Reaction, or PCR). In a quantitative amplification, the amount of amplification product will be proportional to the amount of template in the original sample. Comparison to appropriate controls provides a measure of the amount of prostate cancer-associated RNA. Methods of quantitative amplification are well known to those of skill in the art. Detailed protocols for

quantitative PCR are provided, e.g., in Innis et al., PCR Protocols, A Guide to Methods and Applications (1990).

In some embodiments, a TaqMan based assay is used to measure expression. TaqMan based assays use a fluorogenic oligonucleotide probe that contains a 5' fluorescent dye and a 3' quenching agent. The probe hybridizes to a PCR product, but cannot itself be extended due to a blocking agent at the 3' end. When the PCR product is amplified in subsequent cycles, the 5' nuclease activity of the polymerase, e.g., AmpliTaq, results in the cleavage of the TaqMan probe. This cleavage separates the 5' fluorescent dye and the 3' quenching agent, thereby resulting in an increase in fluorescence as a function of

Other suitable amplification methods include, but are not limited to, ligase chain reaction (LCR) (see Wu & Wallace, Genomics 4:560 (1989), Landegren et al., Science 241:1077 (1988), and Barringer et al., Gene 89:117 (1990), transcription amplification (Kwoh et al., Proc. Natl. Acad. Sci. USA 86:1173 (1989)), self-sustained sequence replication (Guatelli et al., Proc. Nat. Acad. Sci. USA 87:1874 (1990)), dot PCR, and linker adapter PCR, etc.

amplification (see, e.g., literature provided by Perkin-Elmer, e.g., www2.perkin-elmer.com).

## Expression of prostate cancer proteins from nucleic acids

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In a preferred embodiment, prostate cancer nucleic acids, e.g., encoding prostate cancer proteins are used to make a variety of expression vectors to express prostate 20 cancer proteins which can then be used in screening assays, as described below. Expression vectors and recombinant DNA technology are well known to those of skill in the art (see, e.g., Ausubel, supra, and Gene Expression Systems (Fernandez & Hoeffler, eds, 1999)) and are used to express proteins. The expression vectors may be either self-replicating extrachromosomal vectors or vectors which integrate into a host genome. Generally, these 25 expression vectors include transcriptional and translational regulatory nucleic acid operably linked to the nucleic acid encoding the prostate cancer protein. The term "control sequences" refers to DNA sequences used for the expression of an operably linked coding sequence in a particular host organism. Control sequences that are suitable for prokaryotes, e.g., include a promoter, optionally an operator sequence, and a ribosome binding site. Eukaryotic cells are 30 known to utilize promoters, polyadenylation signals, and enhancers.

Nucleic acid is "operably linked" when it is placed into a functional relationship with another nucleic acid sequence. For example, DNA for a presequence or secretory leader is operably linked to DNA for a polypeptide if it is expressed as a preprotein that participates in the secretion of the polypeptide; a promoter or enhancer is operably linked to a coding sequence if it affects the transcription of the sequence; or a ribosome binding site is operably linked to a coding sequence if it is positioned so as to facilitate translation. Generally, "operably linked" means that the DNA sequences being linked are contiguous, and, in the case of a secretory leader, contiguous and in reading phase. However, enhancers do not have to be contiguous. Linking is typically accomplished by ligation at convenient restriction sites. If such sites do not exist, synthetic oligonucleotide adaptors or linkers are used in accordance with conventional practice. Transcriptional and translational regulatory nucleic acid will generally be appropriate to the host cell used to express the prostate cancer protein. Numerous types of appropriate expression vectors, and suitable regulatory sequences are known in the art for a variety of host cells.

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In general, transcriptional and translational regulatory sequences may include, but are not limited to, promoter sequences, ribosomal binding sites, transcriptional start and stop sequences, translational start and stop sequences, and enhancer or activator sequences. In a preferred embodiment, the regulatory sequences include a promoter and transcriptional start and stop sequences.

Promoter sequences encode either constitutive or inducible promoters. The promoters may be either naturally occurring promoters or hybrid promoters. Hybrid promoters, which combine elements of more than one promoter, are also known in the art, and are useful in the present invention.

In addition, an expression vector may comprise additional elements. For example, the expression vector may have two replication systems, thus allowing it to be maintained in two organisms, e.g. in mammalian or insect cells for expression and in a procaryotic host for cloning and amplification. Furthermore, for integrating expression vectors, the expression vector contains at least one sequence homologous to the host cell genome, and preferably two homologous sequences which flank the expression construct. The integrating vector may be directed to a specific locus in the host cell by selecting the

appropriate homologous sequence for inclusion in the vector. Constructs for integrating vectors are well known in the art (e.g., Fernandez & Hoeffler, *supra*).

In addition, in a preferred embodiment, the expression vector contains a selectable marker gene to allow the selection of transformed host cells. Selection genes are well known in the art and will vary with the host cell used.

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The prostate cancer proteins of the present invention are produced by culturing a host cell transformed with an expression vector containing nucleic acid encoding a prostate cancer protein, under the appropriate conditions to induce or cause expression of the prostate cancer protein. Conditions appropriate for prostate cancer protein expression will vary with the choice of the expression vector and the host cell, and will be easily ascertained by one skilled in the art through routine experimentation or optimization. For example, the use of constitutive promoters in the expression vector will require optimizing the growth and proliferation of the host cell, while the use of an inducible promoter requires the appropriate growth conditions for induction. In addition, in some embodiments, the timing of the harvest is important. For example, the baculoviral systems used in insect cell expression are lytic viruses, and thus harvest time selection can be crucial for product yield.

Appropriate host cells include yeast, bacteria, archaebacteria, fungi, and insect and animal cells, including mammalian cells. Of particular interest are Saccharomyces cerevisiae and other yeasts, E. coli, Bacillus subtilis, Sf9 cells, C129 cells, 293 cells, Neurospora, BHK, CHO, COS, HeLa cells, HUVEC (human umbilical vein endothelial cells). THP1 cells (a macrophage cell line) and various other human cells and cell lines.

In a preferred embodiment, the prostate cancer proteins are expressed in mammalian cells. Mammalian expression systems are also known in the art, and include retroviral and adenoviral systems. One expression vector system is a retroviral vector system such as is generally described in PCT/US97/01019 and PCT/US97/01048, both of which are hereby expressly incorporated by reference. Of particular use as mammalian promoters are the promoters from mammalian viral genes, since the viral genes are often highly expressed and have a broad host range. Examples include the SV40 early promoter, mouse mammary tumor virus LTR promoter, adenovirus major late promoter, herpes simplex virus promoter, and the CMV promoter (see, e.g., Fernandez & Hoeffler, supra). Typically, transcription termination and polyadenylation sequences recognized by mammalian cells are regulatory

regions located 3' to the translation stop codon and thus, together with the promoter elements, flank the coding sequence. Examples of transcription terminator and polyadenlyation signals include those derived form SV40.

The methods of introducing exogenous nucleic acid into mammalian hosts, as

well as other hosts, is well known in the art, and will vary with the host cell used.

Techniques include dextran-mediated transfection, calcium phosphate precipitation,
polybrene mediated transfection, protoplast fusion, electroporation, viral infection,
encapsulation of the polynucleotide(s) in liposomes, and direct microinjection of the DNA
into nuclei.

In a preferred embodiment, prostate cancer proteins are expressed in bacterial 10 systems. Bacterial expression systems are well known in the art. Promoters from bacteriophage may also be used and are known in the art. In addition, synthetic promoters and hybrid promoters are also useful; e.g., the tac promoter is a hybrid of the trp and lac promoter sequences. Furthermore, a bacterial promoter can include naturally occurring promoters of non-bacterial origin that have the ability to bind bacterial RNA polymerase and 15 initiate transcription. In addition to a functioning promoter sequence, an efficient ribosome binding site is desirable. The expression vector may also include a signal peptide sequence that provides for secretion of the prostate cancer protein in bacteria. The protein is either secreted into the growth media (gram-positive bacteria) or into the periplasmic space, located 20 between the inner and outer membrane of the cell (gram-negative bacteria). The bacterial expression vector may also include a selectable marker gene to allow for the selection of bacterial strains that have been transformed. Suitable selection genes include genes which render the bacteria resistant to drugs such as ampicillin, chloramphenicol, erythromycin, kanamycin, neomycin and tetracycline. Selectable markers also include biosynthetic genes, such as those in the histidine, tryptophan and leucine biosynthetic pathways. These 25 components are assembled into expression vectors. Expression vectors for bacteria are well known in the art, and include vectors for Bacillus subtilis, E. coli, Streptococcus cremoris, and Streptococcus lividans, among others (e.g., Fernandez & Hoeffler, supra). The bacterial expression vectors are transformed into bacterial host cells using techniques well known in the art, such as calcium chloride treatment, electroporation, and others. 30

In one embodiment, prostate cancer proteins are produced in insect cells. Expression vectors for the transformation of insect cells, and in particular, baculovirus-based expression vectors, are well known in the art.

In a preferred embodiment, prostate cancer protein is produced in yeast cells. Yeast expression systems are well known in the art, and include expression vectors for 5 Saccharomyces cerevisiae, Candida albicans and C. maltosa, Hansenula polymorpha, Kluvveromyces fragilis and K. lactis, Pichia guillerimondii and P. pastoris, Schizosaccharomyces pombe, and Yarrowia lipolytica.

The prostate cancer protein may also be made as a fusion protein, using techniques well known in the art. Thus, e.g., for the creation of monoclonal antibodies, if the 10 desired epitope is small, the prostate cancer protein may be fused to a carrier protein to form an immunogen. Alternatively, the prostate cancer protein may be made as a fusion protein to increase expression, or for other reasons. For example, when the prostate cancer protein is a prostate cancer peptide, the nucleic acid encoding the peptide may be linked to other nucleic acid for expression purposes.

In a preferred embodiment, the prostate cancer protein is purified or isolated after expression. Prostate cancer proteins may be isolated or purified in a variety of ways known to those skilled in the art depending on what other components are present in the sample. Standard purification methods include electrophoretic, molecular, immunological and chromatographic techniques, including ion exchange, hydrophobic, affinity, and reversephase HPLC chromatography, and chromatofocusing. For example, the prostate cancer protein may be purified using a standard anti-prostate cancer protein antibody column. Ultrafiltration and diafiltration techniques, in conjunction with protein concentration, are also useful. For general guidance in suitable purification techniques, see Scopes, Protein Purification (1982). The degree of purification necessary will vary depending on the use of the prostate cancer protein. In some instances no purification will be necessary.

Once expressed and purified if necessary, the prostate cancer proteins and nucleic acids are useful in a number of applications. They may be used as immunoselection reagents, as vaccine reagents, as screening agents, etc.

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### Variants of prostate cancer proteins

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In one embodiment, the prostate cancer proteins are derivative or variant prostate cancer proteins as compared to the wild-type sequence. That is, as outlined more fully below, the derivative prostate cancer peptide will often contain at least one amino acid substitution, deletion or insertion, with amino acid substitutions being particularly preferred. The amino acid substitution, insertion or deletion may occur at any residue within the prostate cancer peptide.

Also included within one embodiment of prostate cancer proteins of the present invention are amino acid sequence variants. These variants typically fall into one or more of three classes: substitutional, insertional or deletional variants. These variants ordinarily are prepared by site specific mutagenesis of nucleotides in the DNA encoding the prostate cancer protein, using cassette or PCR mutagenesis or other techniques well known in the art, to produce DNA encoding the variant, and thereafter expressing the DNA in recombinant cell culture as outlined above. However, variant prostate cancer protein fragments having up to about 100-150 residues may be prepared by in vitro synthesis using established techniques. Amino acid sequence variants are characterized by the predetermined nature of the variation, a feature that sets them apart from naturally occurring allelic or interspecies variation of the prostate cancer protein amino acid sequence. The variants typically exhibit the same qualitative biological activity as the naturally occurring analogue, 20 although variants can also be selected which have modified characteristics as will be more fully outlined below.

While the site or region for introducing an amino acid sequence variation is predetermined, the mutation per se need not be predetermined. For example, in order to optimize the performance of a mutation at a given site, random mutagenesis may be conducted at the target codon or region and the expressed prostate cancer variants screened for the optimal combination of desired activity. Techniques for making substitution mutations at predetermined sites in DNA having a known sequence are well known, e.g., M13 primer mutagenesis and PCR mutagenesis. Screening of the mutants is done using assays of prostate cancer protein activities.

Amino acid substitutions are typically of single residues; insertions usually will be on the order of from about 1 to 20 amino acids, although considerably larger

insertions may be tolerated. Deletions range from about 1 to about 20 residues, although in some cases deletions may be much larger.

Substitutions, deletions, insertions or any combination thereof may be used to arrive at a final derivative. Generally these changes are done on a few amino acids to minimize the alteration of the molecule. However, larger changes may be tolerated in certain circumstances. When small alterations in the characteristics of the prostate cancer protein are desired, substitutions are generally made in accordance with the amino acid substitution relationships provided in the definition section.

The variants typically exhibit the same qualitative biological activity and will

elicit the same immune response as the naturally-occurring analog, although variants also are
selected to modify the characteristics of the prostate cancer proteins as needed. Alternatively,
the variant may be designed such that the biological activity of the prostate cancer protein is
altered. For example, glycosylation sites may be altered or removed.

Substantial changes in function or immunological identity are made by selecting substitutions that are less conservative than those described above. For example, substitutions may be made which more significantly affect: the structure of the polypeptide backbone in the area of the alteration, for example the alpha-helical or beta-sheet structure; the charge or hydrophobicity of the molecule at the target site; or the bulk of the side chain. The substitutions which in general are expected to produce the greatest changes in the polypeptide's properties are those in which (a) a hydrophilic residue, e.g. seryl or threonyl is substituted for (or by) a hydrophobic residue, e.g. leucyl, isoleucyl, phenylalanyl, valyl or alanyl; (b) a cysteine or proline is substituted for (or by) any other residue; (c) a residue having an electropositive side chain, e.g. lysyl, arginyl, or histidyl, is substituted for (or by) an electronegative residue, e.g. glutamyl or aspartyl; or (d) a residue having a bulky side chain, e.g. phenylalanie, is substituted for (or by) one not having a side chain, e.g. glycine.

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Covalent modifications of prostate cancer polypeptides are included within the scope of this invention. One type of covalent modification includes reacting targeted amino acid residues of a prostate cancer polypeptide with an organic derivatizing agent that is capable of reacting with selected side chains or the N-or C-terminal residues of a prostate cancer polypeptide. Derivatization with bifunctional agents is useful, for instance, for crosslinking prostate cancer polypeptides to a water-insoluble support matrix or surface for

use in the method for purifying anti-prostate cancer polypeptide antibodies or screening assays, as is more fully described below. Commonly used crosslinking agents include, e.g., 1,1-bis(diazoacetyl)-2-phenylethane, glutaraldehyde, N-hydroxysuccinimide esters, e.g., esters with 4-azidosalicylic acid, homobifunctional imidoesters, including disuccinimidyl esters such as 3,3'-dithiobis(succinimidylpropionate), bifunctional maleimides such as bis-N-maleimido-1,8-octane and agents such as methyl-3-((p-azidophenyl)dithio)propioimidate.

Other modifications include deamidation of glutaminyl and asparaginyl residues to the corresponding glutamyl and aspartyl residues, respectively, hydroxylation of proline and lysine, phosphorylation of hydroxyl groups of seryl, threonyl or tyrosyl residues, methylation of the amino groups of the lysine, arginine, and histidine side chains (Creighton, Proteins: Structure and Molecular Properties, pp. 79-86 (1983)), acetylation of the Neterminal amine, and amidation of any C-terminal carboxyl group.

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Another type of covalent modification of the prostate cancer polypeptide included within the scope of this invention comprises altering the native glycosylation pattern of the polypeptide. "Altering the native glycosylation pattern" is intended for purposes herein to mean deleting one or more carbohydrate moieties found in native sequence prostate cancer polypeptide, and/or adding one or more glycosylation sites that are not present in the native sequence prostate cancer polypeptide. Glycosylation patterns can be altered in many ways. For example the use of different cell types to express prostate cancer-associated sequences can result in different glycosylation patterns.

Addition of glycosylation sites to prostate cancer polypeptides may also be accomplished by altering the amino acid sequence thereof. The alteration may be made, e.g., by the addition of, or substitution by, one or more serine or threonine residues to the native sequence prostate cancer polypeptide (for O-linked glycosylation sites). The prostate cancer amino acid sequence may optionally be altered through changes at the DNA level, particularly by mutating the DNA encoding the prostate cancer polypeptide at preselected bases such that codons are generated that will translate into the desired amino acids.

Another means of increasing the number of carbohydrate moieties on the prostate cancer polypeptide is by chemical or enzymatic coupling of glycosides to the polypeptide. Such methods are described in the art, e.g., in WO 87/05330, and in Aplin & Wriston, CRC Crit. Rev. Biochem., pp. 259-306 (1981).

Removal of carbohydrate moieties present on the prostate cancer polypeptide may be accomplished chemically or enzymatically or by mutational substitution of codons encoding for amino acid residues that serve as targets for glycosylation. Chemical deglycosylation techniques are known in the art and described, for instance, by Hakimuddin, et al., Arch. Biochem. Biophys., 259:52 (1987) and by Edge et al., Anal. Biochem., 118:131 (1981). Enzymatic cleavage of carbohydrate moieties on polypeptides can be achieved by the use of a variety of endo-and exo-glycosidases as described by Thotakura et al., Meth. Enzymol., 138:350 (1987).

Another type of covalent modification of prostate cancer comprises linking the prostate cancer polypeptide to one of a variety of nonproteinaceous polymers, e.g., polyethylene glycol, polypropylene glycol, or polyoxyalkylenes, in the manner set forth in U.S. Patent Nos. 4,640,835; 4,496,689; 4,301,144; 4,670,417; 4,791,192 or 4,179,337.

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Prostate cancer polypeptides of the present invention may also be modified in a way to form chimeric molecules comprising a prostate cancer polypeptide fused to another, heterologous polypeptide or amino acid sequence. In one embodiment, such a chimeric molecule comprises a fusion of a prostate cancer polypeptide with a tag polypeptide which provides an epitope to which an anti-tag antibody can selectively bind. The epitope tag is generally placed at the amino-or carboxyl-terminus of the prostate cancer polypeptide. The presence of such epitope-tagged forms of a prostate cancer polypeptide can be deteeted using an antibody against the tag polypeptide. Also, provision of the epitope tag enables the prostate cancer polypeptide to be readily purified by affinity purification using an anti-tag antibody or another type of affinity matrix that binds to the epitope tag. In an alternative embodiment, the chimeric molecule may comprise a fusion of a prostate cancer polypeptide with an immunoglobulin or a particular region of an immunoglobulin. For a bivalent form of the chimeric molecule, such a fusion could be to the Fe region of an IgG molecule.

Various tag polypeptides and their respective antibodies are well known in the art. Examples include poly-histidine (poly-his) or poly-histidine-glycine (poly-his-gly) tags; HIS6 and metal chelation tags, the flu HA tag polypeptide and its antibody 12CA5 (Field et al., Mol. Cell. Biol. 8:2159-2165 (1988)); the c-myc tag and the 8F9, 3C7, 6E10, G4, B7 and 9E10 antibodies thereto (Evan et al., Molecular and Cellular Biology 5:3610-3616 (1985)); and the Herpes Simplex virus glycoprotein D (gD) tag and its antibody (Paborsky et al.,

Protein Engineering 3(6):547-553 (1990)). Other tag polypeptides include the Flag-peptide (Hopp et al., BioTechnology 6:1204-1210 (1988)); the KT3 epitope peptide (Martin et al., Science 255:192-194 (1992)); tubulin epitope peptide (Skinner et al., J. Biol. Chem. 266:15163-15166 (1991)); and the T7 gene 10 protein peptide tag (Lutz-Freyermuth et al., Proc. Natl. Acad. Sci. USA 87:6393-6397 (1990)).

Also included are other prostate cancer proteins of the prostate cancer family, and prostate cancer proteins from other organisms, which are cloned and expressed as outlined below. Thus, probe or degenerate polymerase chain reaction (PCR) primer sequences may be used to find other related prostate cancer proteins from humans or other organisms. As will be appreciated by those in the art, particularly useful probe and/or PCR primer sequences include the unique areas of the prostate cancer nucleic acid sequence. As is generally known in the art, preferred PCR primers are from about 15 to about 35 nucleotides in length, with from about 20 to about 30 being preferred, and may contain inosine as needed. The conditions for the PCR reaction are well known in the art (e.g., Innis, PCR Protocols, supra).

### Antibodies to prostate cancer proteins

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In a preferred embodiment, when the prostate cancer protein is to be used to generate antibodies, e.g., for immunotherapy or immunodiagnosis, the prostate cancer protein should share at least one epitope or determinant with the full length protein. By "epitope" or "determinant" herein is typically meant a portion of a protein which will generate and/or bind an antibody or T-cell receptor in the context of MHC. Thus, in most instances, antibodies made to a smaller prostate cancer protein will be able to bind to the full-length protein, particularly linear epitopes. In a preferred embodiment; the epitope is unique; that is, antibodies generated to a unique epitope show little or no cross-reactivity.

Methods of preparing polyclonal antibodies are known to the skilled artisan (e.g., Coligan, supra; and Harlow & Lane, supra). Polyclonal antibodies can be raised in a mammal, e.g., by one or more injections of an immunizing agent and, if desired, an adjuvant. Typically, the immunizing agent and/or adjuvant will be injected in the mammal by multiple subcutaneous or intraperitoneal injections. The immunizing agent may include a protein encoded by a nucleic acid of the figures or fragment thereof or a fusion protein thereof. It

may be useful to conjugate the immunizing agent to a protein known to be immunogenic in the mammal being immunized. Examples of such immunogenic proteins include but are not limited to keyhole limpet hemocyanin, serum albumin, bovine thyroglobulin, and soybean trypsin inhibitor. Examples of adjuvants which may be employed include Freund's complete adjuvant and MPL-TDM adjuvant (monophosphoryl Lipid A, synthetic trehalose dicorynomycolate). The immunization protocol may be selected by one skilled in the art without undue experimentation.

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The antibodies may, alternatively, be monoclonal antibodies. Monoclonal antibodies may be prepared using hybridoma methods, such as those described by Kohler & Milstein, Nature 256:495 (1975). In a hybridoma method, a mouse, hamster, or other appropriate host animal, is typically immunized with an immunizing agent to elicit lymphocytes that produce or are capable of producing antibodies that will specifically bind to the immunizing agent. Alternatively, the lymphocytes may be immunized in vitro. The immunizing agent will typically include a polypeptide encoded by a nucleic acid of Tables 1-16 fragment thereof, or a fusion protein thereof. Generally, either peripheral blood lymphocytes ("PBLs") are used if cells of human origin are desired, or spleen cells or lymph node cells are used if non-human mammalian sources are desired. The lymphocytes are then fused with an immortalized cell line using a suitable fusing agent, such as polyethylene glycol, to form a hybridoma cell (Goding, Monoclonal Antibodies: Principles and Practice, pp. 59-103 (1986)). Immortalized cell lines are usually transformed mammalian cells, particularly myeloma cells of rodent, bovine and human origin. Usually, rat or mouse myeloma cell lines are employed. The hybridoma cells may be cultured in a suitable culture medium that preferably contains one or more substances that inhibit the growth or survival of the unfused, immortalized cells. For example, if the parental cells lack the enzyme hypoxanthine guanine phosphoribosyl transferase (HGPRT or HPRT), the culture medium for the hybridomas typically will include hypoxanthine, aminopterin, and thymidine ("HAT

In one embodiment, the antibodies are bispecific antibodies. Bispecific antibodies are monoclonal, preferably human or humanized, antibodies that have binding specificities for at least two different antigens or that have binding specificities for two epitopes on the same antigen. In one embodiment, one of the binding specificities is for a

medium"), which substances prevent the growth of HGPRT-deficient cells.

protein encoded by a nucleic acid Tables 1-16 or a fragment thereof, the other one is for any other antigen, and preferably for a cell-surface protein or receptor or receptor subunit, preferably one that is tumor specific. Alternatively, tetramer-type technology may create multivalent reagents.

In a preferred embodiment, the antibodies to prostate cancer protein are capable of reducing or eliminating a biological function of a prostate cancer protein, as is described below. That is, the addition of anti-prostate cancer protein antibodies (either polyclonal or preferably monoclonal) to prostate cancer tissue (or cells containing prostate cancer) may reduce or eliminate the prostate cancer. Generally, at least a 25% decrease in activity, growth, size or the like is preferred, with at least about 50% being particularly preferred and about a 95-100% decrease being especially preferred.

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In a preferred embodiment the antibodies to the prostate cancer proteins are humanized antibodies (e.g., Xenerex Biosciences, Mederex, Inc., Abgenix, Inc., Protein Design Labs.Inc.) Humanized forms of non-human (e.g., murine) antibodies are chimeric molecules of immunoglobulins, immunoglobulin chains or fragments thereof (such as Fv, Fab, Fab', F(ab')2 or other antigen-binding subsequences of antibodies) which contain minimal sequence derived from non-human immunoglobulin. Humanized antibodies include human immunoglobulins (recipient antibody) in which residues from a complementary determining region (CDR) of the recipient are replaced by residues from a CDR of a nonhuman species (donor antibody) such as mouse, rat or rabbit having the desired specificity, affinity and capacity. In some instances, Fv framework residues of the human immunoglobulin are replaced by corresponding non-human residues. Humanized antibodies may also comprise residues which are found neither in the recipient antibody nor in the imported CDR or framework sequences. In general, a humanized antibody will comprise substantially all of at least one, and typically two, variable domains, in which all or substantially all of the CDR regions correspond to those of a non-human immunoglobulin and all or substantially all of the framework (FR) regions are those of a human immunoglobulin consensus sequence. The humanized antibody optimally also will comprise at least a portion of an immunoglobulin constant region (Fc), typically that of a human immunoglobulin (Jones et al., Nature 321:522-525 (1986); Riechmann et al., Nature 332:323-329 (1988); and Presta, Curr. Op. Struct. Biol. 2:593-596 (1992)). Humanization

can be essentially performed following the method of Winter and co-workers (Jones et al., Nature 321:522-525 (1986); Riechmann et al., Nature 332:323-327 (1988); Verhoeyen et al., Science 239:1534-1536 (1988)), by substituting rodent CDRs or CDR sequences for the corresponding sequences of a human antibody. Accordingly, such humanized antibodies are chimeric antibodies (U.S. Patent No. 4,816,567), wherein substantially less than an intact human variable domain has been substituted by the corresponding sequence from a non-human species.

Human antibodies can also be produced using various techniques known in the art, including phage display libraries (Hoogenboom & Winter, J. Mol. Biol. 227:381 (1991); 10 Marks et al., J. Mol. Biol. 222:581 (1991)). The techniques of Cole et al. and Boerner et al. are also available for the preparation of human monoclonal antibodies (Cole et al., Monoclonal Antibodies and Cancer Therapy, p. 77 (1985) and Boerner et al., J. Immunol. 147(1):86-95 (1991)). Similarly, human antibodies can be made by introducing of human immunoglobulin loci into transgenic animals, e.g., mice in which the endogenous immunoglobulin genes have been partially or completely inactivated. Upon challenge. 15 human antibody production is observed, which closely resembles that seen in humans in all respects, including gene rearrangement, assembly, and antibody repertoire. This approach is described, e.g., in U.S. Patent Nos. 5,545,807; 5,545,806; 5,569,825; 5,625,126; 5,633,425; 5.661.016, and in the following scientific publications: Marks et al., Bio/Technology 10:779-783 (1992); Lonberg et al., Nature 368:856-859 (1994); Morrison, Nature 368:812-13 (1994); Fishwild et al., Nature Biotechnology 14:845-51 (1996); Neuberger, Nature Biotechnology 14:826 (1996); Lonberg & Huszar, Intern. Rev. Immunol. 13:65-93 (1995). By immunotherapy is meant treatment of prostate cancer with an antibody raised against prostate cancer proteins. As used herein, immunotherapy can be passive or active. Passive immunotherapy as defined herein is the passive transfer of antibody to a 25 recipient (patient). Active immunization is the induction of antibody and/or T-cell responses in a recipient (patient). Induction of an immune response is the result of providing the recipient with an antigen to which antibodies are raised. As appreciated by one of ordinary skill in the art, the antigen may be provided by injecting a polypeptide against which 30 antibodies are desired to be raised into a recipient, or contacting the recipient with a nucleic

acid capable of expressing the antigen and under conditions for expression of the antigen, leading to an immune response.

In a preferred embodiment the prostate cancer proteins against which antibodies are raised are secreted proteins as described above. Without being bound by theory, antibodies used for treatment, bind and prevent the secreted protein from binding to its receptor, thereby inactivating the secreted prostate cancer protein.

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In another preferred embodiment, the prostate cancer protein to which antibodies are raised is a transmembrane protein. Without being bound by theory, antibodies used for treatment, bind the extracellular domain of the prostate cancer protein and prevent it from binding to other proteins, such as circulating ligands or cell-associated molecules. The antibody may cause down-regulation of the transmembrane prostate cancer protein. As will be appreciated by one of ordinary skill in the art, the antibody may be a competitive, noncompetitive or uncompetitive inhibitor of protein binding to the extracellular domain of the prostate cancer protein. The antibody is also an antagonist of the prostate cancer protein. Further, the antibody prevents activation of the transmembrane prostate cancer protein. In one aspect, when the antibody prevents the binding of other molecules to the prostate cancer protein, the antibody prevents growth of the cell. The antibody may also be used to target or sensitize the cell to cytotoxic agents, including, but not limited to TNF-α, TNF-β, IL-1, INF-γ and IL-2, or chemotherapeutic agents including 5FU, vinblastine, actinomycin D, cisplatin, methotrexate, and the like. In some instances the antibody belongs to a sub-type that activates serum complement when complexed with the transmembrane protein thereby mediating cytotoxicity or antigen-dependent cytotoxicity (ADCC). Thus, prostate cancer is treated by administering to a patient antibodies directed against the transmembrane prostate caneer protein. Antibody-labeling may activate a co-toxin, localize a toxin payload, or

In another preferred embodiment, the antibody is conjugated to an effector moiety. The effector moiety can be any number of molecules, including labelling moieties such as radioactive labels or fluorescent labels, or can be a therapeutic moiety. In one aspect the therapeutic moiety is a small molecule that modulates the activity of the prostate cancer protein. In another aspect the therapeutic moiety modulates the activity of molecules associated with or in close proximity to the prostate cancer protein. The therapeutic moiety

otherwise provide means to locally ablate cells.

may inhibit enzymatic activity such as protease or collagenase or protein kinase activity associated with prostate cancer.

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In a preferred embodiment, the therapeutic moiety can also be a cytotoxic agent. In this method, targeting the cytotoxic agent to prostate cancer tissue or cells, results in a reduction in the number of afflicted cells, thereby reducing symptoms associated with prostate cancer. Cytotoxic agents are numerous and varied and include, but are not limited to, cytotoxic drugs or toxins or active fragments of such toxins. Suitable toxins and their corresponding fragments include diphtheria A chain, exotoxin A chain, ricin A chain, abrin A chain, curcin, crotin, phenomycin, enomycin and the like. Cytotoxic agents also include radiochemicals made by conjugating radioisotopes to antibodies raised against prostate cancer proteins, or binding of a radionuclide to a chelating agent that has been covalently attached to the antibody. Targeting the therapeutic moiety to transmembrane prostate cancer proteins not only serves to increase the local concentration of therapeutic moiety in the prostate cancer afflicted area, but also serves to reduce deleterious side effects that may be associated with the therapeutic moiety.

In another preferred embodiment, the prostate cancer protein against which the antibodies are raised is an intracellular protein. In this case, the antibody may be conjugated to a protein which facilitates entry into the cell. In one case, the antibody enters the cell by endocytosis. In another embodiment, a nucleic acid encoding the antibody is administered to the individual or cell. Moreover, wherein the prostate cancer protein can be targeted within a cell, i.e., the nucleus, an antibody thereto contains a signal for that target localization, i.e., a nuclear localization signal.

The prostate cancer antibodies of the invention specifically bind to prostate cancer proteins. By "specifically bind" herein is meant that the antibodies bind to the protein with a  $K_d$  of at least about 0.1 mM, more usually at least about 1  $\mu M$ , preferably at least about 0.1  $\mu M$  or better, and most preferably, 0.01  $\mu M$  or better. Selectivity of binding is also important.

# Detection of prostate cancer sequence for diagnostic and therapeutic applications

In one aspect, the RNA expression levels of genes are determined for different cellular states in the prostate cancer phenotype. Expression levels of genes in normal tissue

(i.e., not undergoing prostate cancer) and in prostate cancer tissue (and in some cases, for varying severities of prostate cancer that relate to prognosis, as outlined below) are evaluated to provide expression profiles. An expression profile of a particular cell state or point of development is essentially a "fingerprint" of the state. While two states may have any particular gene similarly expressed, the evaluation of a number of genes simultaneously allows the generation of a gene expression profile that is reflective of the state of the cell. By comparing expression profiles of cells in different states, information regarding which genes are important (including both up- and down-regulation of genes) in each of these states is obtained. Then, diagnosis may be performed or confirmed to determine whether a tissue sample has the gene expression profile of normal or cancerous tissue. This will provide for molecular diagnosis of related conditions.

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"Differential expression," or grammatical equivalents as used herein, refers to qualitative or quantitative differences in the temporal and/or cellular gene expression patterns within and among cells and tissue. Thus, a differentially expressed gene can qualitatively have its expression altered, including an activation or inactivation, in, e.g., normal versus prostate cancer tissue. Genes may be turned on or turned off in a particular state, relative to another state thus permitting comparison of two or more states. A qualitatively regulated gene will exhibit an expression pattern within a state or cell type which is detectable by standard techniques. Some genes will be expressed in one state or cell type, but not in both. Alternatively, the difference in expression may be quantitative, e.g., in that expression is increased or decreased; i.e., gene expression is either upregulated, resulting in an increased amount of transcript, or downregulated, resulting in a decreased amount of transcript. The degree to which expression differs need only be large enough to quantify via standard characterization techniques as outlined below, such as by use of Affymetrix GeneChip™ expression arrays, Lockhart, Nature Biotechnology 14:1675-1680 (1996), 25 hereby expressly incorporated by reference. Other techniques include, but are not limited to, quantitative reverse transcriptase PCR, northern analysis and RNase protection. As outlined above, preferably the change in expression (i.e., upregulation or downregulation) is at least about 50%, more preferably at least about 100%, more preferably at least about 150%, more preferably at least about 200%, with from 300 to at least 1000% being especially preferred.

Evaluation may be at the gene transcript, or the protein level. The amount of gene expression may be monitored using nucleic acid probes to the DNA or RNA equivalent of the gene transcript, and the quantification of gene expression levels, or, alternatively, the final gene product itself (protein) can be monitored, e.g., with antibodies to the prostate cancer protein and standard immunoassays (ELISAs, etc.) or other techniques, including mass spectroscopy assays, 2D gel electrophoresis assays, etc. Proteins corresponding to prostate cancer genes, i.e., those identified as being important in a prostate cancer phenotype, can be evaluated in a prostate cancer diagnostic test.

In a preferred embodiment, gene expression monitoring is performed simultaneously on a number of genes. Multiple protein expression monitoring can be performed as well. Similarly, these assays may be performed on an individual basis as well.

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In this embodiment, the prostate cancer nucleic acid probes are attached to biochips as outlined herein for the detection and quantification of prostate cancer sequences in a particular cell. The assays are further described below in the example. PCR techniques can be used to provide greater sensitivity.

In a preferred embodiment nucleic acids encoding the prostate cancer protein are detected. Although DNA or RNA encoding the prostate cancer protein may be detected, of particular interest are methods wherein an mRNA encoding a prostate cancer protein is detected. Probes to detect mRNA can be a nucleotide/deoxynucleotide probe that is complementary to and hybridizes with the mRNA and includes, but is not limited to, oligonucleotides, cDNA or RNA. Probes also should contain a detectable label, as defined herein. In one method the mRNA is detected after immobilizing the nucleic acid to be examined on a solid support such as nylon membranes and hybridizing the probe with the sample. Following washing to remove the non-specifically bound probe, the label is detected. In another method detection of the mRNA is performed in situ. In this method permeabilized cells or tissue samples are contacted with a detectably labeled nucleic acid probe for sufficient time to allow the probe to hybridize with the target mRNA. Following washing to remove the non-specifically bound probe, the label is detected. For example a digoxygenin labeled riboprobe (RNA probe) that is complementary to the mRNA encoding a prostate cancer protein is detected by binding the digoxygenin with an anti-digoxygenin

secondary antibody and developed with nitro blue tetrazolium and 5-bromo-4-chloro-3indovl phosphate.

In a preferred embodiment, various proteins from the three classes of proteins as described herein (secreted, transmembrane or intracellular proteins) are used in diagnostic assays. The prostate cancer proteins, antibodies, nucleic acids, modified proteins and cells containing prostate cancer sequences are used in diagnostic assays. This can be performed on an individual gene or corresponding polypeptide level. In a preferred embodiment, the expression profiles are used, preferably in conjunction with high throughput screening techniques to allow monitoring for expression profile genes and/or corresponding polypeptides.

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As described and defined herein, prostate cancer proteins, including intracellular, transmembrane or secreted proteins, find use as markers of prostate cancer. Detection of these proteins in putative prostate cancer tissue allows for detection or diagnosis of prostate cancer. In one embodiment, antibodies are used to detect prostate cancer proteins. A preferred method separates proteins from a sample by electrophoresis on a gel (typically a denaturing and reducing protein gel, but may be another type of gel, including isoelectric focusing gels and the like). Following separation of proteins, the prostate cancer protein is detected, e.g., by immunoblotting with antibodies raised against the prostate cancer protein. Methods of immunoblotting are well known to those of ordinary skill in the art.

In another preferred method, antibodies to the prostate cancer protein find use in in situ imaging techniques, e.g., in histology (e.g., Methods in Cell Biology: Antibodies in Cell Biology, volume 37 (Asai, ed. 1993)). In this method cells are contacted with from one to many antibodies to the prostate cancer protein(s). Following washing to remove nonspecific antibody binding, the presence of the antibody or antibodies is detected. In one embodiment the antibody is detected by incubating with a secondary antibody that contains a detectable label. In another method the primary antibody to the prostate cancer protein(s) contains a detectable label, e.g. an enzyme marker that can act on a substrate. In another preferred embodiment each one of multiple primary antibodies contains a distinct and detectable label. This method finds particular use in simultaneous screening for a plurality of prostate cancer proteins. As will be appreciated by one of ordinary skill in the art, many other histological imaging techniques are also provided by the invention.

In a preferred embodiment the label is detected in a fluorometer which has the ability to detect and distinguish emissions of different wavelengths. In addition, a fluorescence activated cell sorter (FACS) can be used in the method.

In another preferred embodiment, antibodies find use in diagnosing prostate

cancer from blood, serum, plasma, stool, and other samples. Such samples, therefore, are
useful as samples to be probed or tested for the presence of prostate cancer proteins.

Antibodies can be used to detect a prostate cancer protein by previously described
immunoassay techniques including ELISA, immunoblotting (western blotting),
immunoprecipitation, BIACORE technology and the like. Conversely, the presence of
antibodies may indicate an immune response against an endogenous prostate cancer protein.

In a preferred embodiment, in situ hybridization of labeled prostate cancer nucleic acid probes to tissue arrays is done. For example, arrays of tissue samples, including prostate cancer tissue and/or normal tissue, are made. In situ hybridization (see, e.g., Ausubel, supra) is then performed. When comparing the fingerprints between an individual and a standard, the skilled artisan can make a diagnosis, a prognosis, or a prediction based on the findings. It is further understood that the genes which indicate the diagnosis may differ from those which indicate the prognosis and molecular profiling of the condition of the cells may lead to distinctions between responsive or refractory conditions or may be predictive of outcomes.

In a preferred embodiment, the prostate cancer proteins, antibodies, nucleic acids, modified proteins and cells containing prostate cancer sequences are used in prognosis assays. As above, gene expression profiles can be generated that correlate to prostate cancer, in terms of long term prognosis. Again, this may be done on either a protein or gene level, with the use of genes being preferred. As above, prostate cancer probes may be attached to biochips for the detection and quantification of prostate cancer sequences in a tissue or patient. The assays proceed as outlined above for diagnosis. PCR method may provide more sensitive and accurate quantification.

#### Assays for therapeutic compounds

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In a preferred embodiment members of the proteins, nucleic acids, and antibodies as described herein are used in drug screening assays. The prostate cancer

proteins, antibodies, nucleic acids, modified proteins and cells containing prostate cancer sequences are used in drug screening assays or by evaluating the effect of drug candidates on a "gene expression profile" or expression profile of polypeptides. In a preferred embodiment, the expression profiles are used, preferably in conjunction with high throughput screening techniques to allow monitoring for expression profile genes after treatment with a candidate agent (e.g., Zlokarnik, et al., Science 279:84-8 (1998); Heid, Genome Res 6:986-94, 1996).

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In a preferred embodiment, the prostate cancer proteins, antibodies, nucleic acids, modified proteins and cells containing the native or modified prostate cancer proteins are used in screening assays. That is, the present invention provides novel methods for screening for compositions which modulate the prostate cancer phenotype or an identified physiological function of a prostate cancer protein. As above, this can be done on an individual gene level or by evaluating the effect of drug candidates on a "gene expression profile". In a preferred embodiment, the expression profiles are used, preferably in conjunction with high throughput screening techniques to allow monitoring for expression profile genes after treatment with a candidate agent, see Zlokamik, supra.

Having identified the differentially expressed genes herein, a variety of assays may be executed. In a preferred embodiment, assays may be run on an individual gene or protein level. That is, having identified a particular gene as up regulated in prostate cancer, test compounds can be screened for the ability to modulate gene expression or for binding to the prostate cancer protein. "Modulation" thus includes both an increase and a decrease in gene expression. The preferred amount of modulation will depend on the original change of the gene expression in normal versus tissue undergoing prostate cancer, with changes of at least 10%, preferably 50%, more preferably 100-300%, and in some embodiments 300-1000% or greater. Thus, if a gene exhibits a 4-fold increase in prostate cancer tissue compared to normal tissue, a decrease of about four-fold is often desired; similarly, a 10-fold decrease in prostate cancer tissue compared to normal tissue often provides a target value of a 10-fold increase in expression to be induced by the test compound.

The amount of gene expression may be monitored using nucleic acid probes and the quantification of gene expression levels, or, alternatively, the gene product itself can be monitored, e.g., through the use of antibodies to the prostate cancer protein and standard

immunoassays. Proteomics and separation techniques may also allow quantification of expression.

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In a preferred embodiment, gene expression or protein monitoring of a number of entities, i.e., an expression profile, is monitored simultaneously. Such profiles will twoically involve a plurality of those entities described herein..

In this embodiment, the prostate cancer nucleic acid probes are attached to biochips as outlined herein for the detection and quantification of prostate cancer sequences in a particular cell. Alternatively, PCR may be used. Thus, a series, e.g., of microtiter plate, may be used with dispensed primers in desired wells. A PCR reaction can then be performed and analyzed for each well.

Expression monitoring can be performed to identify compounds that modify the expression of one or more prostate cancer-associated sequences, e.g., a polynucleotide sequence set out in Tables 1-16. Generally, in a preferred embodiment, a test modulator is added to the cells prior to analysis. Moreover, screens are also provided to identify agents that modulate prostate cancer, modulate prostate cancer proteins, bind to a prostate cancer protein, or interfere with the binding of a prostate cancer protein and an antibody or other binding partner.

The term "test compound" or "drug candidate" or "modulator" or grammatical equivalents as used herein describes any molecule, e.g., protein, oligopeptide, small organic molecule, polysaccharide, polynucleotide, etc., to be tested for the capacity to directly or indirectly alter the prostate cancer phenotype or the expression of a prostate cancer sequence, e.g., a nucleic acid or protein sequence. In preferred embodiments, modulators alter expression profiles, or expression profile nucleic acids or proteins provided herein. In one embodiment, the modulator suppresses a prostate cancer phenotype, e.g. to a normal tissue fingerprint. In another embodiment, a modulator induced a prostate cancer phenotype. Generally, a plurality of assay mixtures are run in parallel with different agent concentrations to obtain a differential response to the various concentrations. Typically, one of these concentrations serves as a negative control, i.e., at zero concentration or below the level of detection.

Drug candidates encompass numerous chemical classes, though typically they are organic molecules, preferably small organic compounds having a molecular weight of

more than 100 and less than about 2,500 daltons. Preferred small molecules are less than 2000, or less than 1500 or less than 1000 or less than 500 D. Candidate agents comprise functional groups necessary for structural interaction with proteins, particularly hydrogen bonding, and typically include at least an amine, carbonyl, hydroxyl or carboxyl group, preferably at least two of the functional chemical groups. The candidate agents often comprise cyclical carbon or heterocyclic structures and/or aromatic or polyaromatic structures substituted with one or more of the above functional groups. Candidate agents are also found among biomolecules including peptides, saccharides, fatty acids, steroids, purines, pyrimidines, derivatives, structural analogs or combinations thereof. Particularly preferred are peptides.

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In one aspect, a modulator will neutralize the effect of a prostate cancer protein. By "neutralize" is meant that activity of a protein is inhibited or blocked and the consequent effect on the cell.

In certain embodiments, combinatorial libraries of potential modulators will be screened for an ability to bind to a prostate cancer polypeptide or to modulate activity. Conventionally, new chemical entities with useful properties are generated by identifying a chemical compound (called a "lead compound") with some desirable property or activity, e.g., inhibiting activity, creating variants of the lead compound, and evaluating the property and activity of those variant compounds. Often, high throughput screening (HTS) methods are employed for such an analysis.

In one preferred embodiment, high throughput screening methods involve providing a library containing a large number of potential therapeutic compounds (candidate compounds). Such "combinatorial chemical libraries" are then screened in one or more assays to identify those library members (particular chemical species or subclasses) that display a desired characteristic activity. The compounds thus identified can serve as conventional "lead compounds" or can themselves be used as potential or actual therapeutics.

A combinatorial chemical library is a collection of diverse chemical compounds generated by either chemical synthesis or biological synthesis by combining a number of chemical "building blocks" such as reagents. For example, a linear combinatorial chemical library, such as a polypeptide (e.g., mutein) library, is formed by combining a set of chemical building blocks called amino acids in every possible way for a given compound

length (i.e., the number of amino acids in a polypeptide compound). Millions of chemical compounds can be synthesized through such combinatorial mixing of chemical building blocks (Gallop et al., J. Med. Chem. 37(9):1233-1251 (1994)).

Preparation and screening of combinatorial chemical libraries is well known to those of skill in the art. Such combinatorial chemical libraries include, but are not limited to, 5 peptide libraries (see, e.g., U.S. Patent No. 5,010,175, Furka, Pept. Prot. Res. 37:487-493 (1991), Houghton et al., Nature, 354:84-88 (1991)), peptoids (PCT Publication No WO 91/19735), encoded peptides (PCT Publication WO 93/20242), random bio-oligomers (PCT Publication WO 92/00091), benzodiazepines (U.S. Pat. No. 5,288,514), diversomers such as hydantoins, benzodiazepines and dipeptides (Hobbs et al., Proc. Nat. Acad. Sci. USA 10 90:6909-6913 (1993)), vinylogous polypeptides (Hagihara et al., J. Amer. Chem. Soc. 114:6568 (1992)), nonpeptidal peptidomimetics with a Beta-D-Glucose scaffolding (Hirschmann et al., J. Amer. Chem. Soc. 114:9217-9218 (1992)), analogous organic syntheses of small compound libraries (Chen et al., J. Amer. Chem. Soc. 116:2661 (1994)), oligocarbamates (Cho, et al., Science 261:1303 (1993)), and/or peptidyl phosphonates 15 (Campbell et al., J. Org. Chem. 59:658 (1994)). See, generally, Gordon et al., J. Med. Chem. 37:1385 (1994), nucleic acid libraries (see, e.g., Strategene, Corp.), peptide nucleic acid libraries (see, e.g., U.S. Patent 5,539,083), antibody libraries (see, e.g., Vaughn et al., Nature Biotechnology 14(3):309-314 (1996), and PCT/US96/10287), carbohydrate libraries (see, 20 e.g., Liang et al., Science 274:1520-1522 (1996), and U.S. Patent No. 5,593,853), and small organic molecule libraries (see, e.g., benzodiazepines, Baum, C&EN, Jan 18, page 33 (1993); isoprenoids, U.S. Patent No. 5,569,588; thiazolidinones and metathiazanones, U.S. Patent No. 5,549,974; pyrrolidines, U.S. Patent Nos. 5,525,735 and 5,519,134; morpholino compounds.

Devices for the preparation of combinatorial libraries are commercially available (see, e.g., 357 MPS, 390 MPS, Advanced Chem Tech, Louisville KY, Symphony, Rainin, Woburn, MA, 433A Applied Biosystems, Foster City, CA, 9050 Plus, Millipore, Bedford, MA).

U.S. Patent No. 5,506,337; benzodiazepines, U.S. Patent No. 5,288,514; and the like).

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A number of well known robotic systems have also been developed for solution phase chemistries. These systems include automated workstations like the automated synthesis apparatus developed by Takeda Chemical Industries, LTD. (Osaka,

Japan) and many robotic systems utilizing robotic arms (Zymate II, Zymark Corporation, Hopkinton, Mass.; Orca, Hewlett-Packard, Palo Alto, Calif.), which mimic the manual synthetic operations performed by a chemist. Any of the above devices are suitable for use with the present invention. The nature and implementation of modifications to these devices (if any) so that they can operate as discussed herein will be apparent to persons skilled in the relevant art. In addition, numerous combinatorial libraries are themselves commercially available (see, e.g., ComGenex, Princeton, N.J., Asinex, Moscow, Ru, Tripos, Inc., St. Louis, MO, ChemStar, Ltd, Moscow, RU, 3D Pharmaceuticals, Exton, PA, Martek Biosciences, Columbia, MD, etc.).

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The assays to identify modulators are amenable to high throughput screening. Preferred assays thus detect enhancement or inhibition of prostate cancer gene transcription, inhibition or enhancement of polypeptide expression, and inhibition or enhancement of polypeptide activity.

High throughput assays for the presence, absence, quantification, or other properties of particular nucleic acids or protein products are well known to those of skill in the art. Similarly, binding assays and reporter gene assays are similarly well known. Thus, e.g., U.S. Patent No. 5,559,410 discloses high throughput screening methods for proteins, U.S. Patent No. 5,585,639 discloses high throughput screening methods for nucleic acid binding (i.e., in arrays), while U.S. Patent Nos. 5,576,220 and 5,541,061 disclose high throughput methods of screening for ligand/antibody binding.

In addition, high throughput screening systems are commercially available (see, e.g., Zymark Corp., Hopkinton, MA; Air Technical Industries, Mentor, OH; Beckman Instruments, Inc., Fullerton, CA; Precision Systems, Inc., Natick, MA, etc.). These systems typically automate entire procedures, including all sample and reagent pipetting, liquid dispensing, timed incubations, and final readings of the mieroplate in detector(s) appropriate for the assay. These configurable systems provide high throughput and rapid start up as well as a high degree of flexibility and customization. The manufacturers of such systems provide detailed protocols for various high throughput systems. Thus, e.g., Zymark Corp. provides technical bulletins describing screening systems for detecting the modulation of gene transcription, ligand binding, and the like.

In one embodiment, modulators are proteins, often naturally occurring proteins or fragments of naturally occurring proteins. Thus, e.g., cellular extracts containing proteins, or random or directed digests of proteinaceous cellular extracts, may be used. In this way libraries of proteins may be made for screening in the methods of the invention. Particularly preferred in this embodiment are libraries of bacterial, fungal, viral, and mammalian proteins, with the latter being preferred, and human proteins being especially preferred. Particularly useful test compound will be directed to the class of proteins to which the target belongs, e.g., substrates for enzymes or ligands and receptors.

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In a preferred embodiment, modulators are peptides of from about 5 to about 30 amino acids, with from about 5 to about 20 amino acids being preferred, and from about 7 to about 15 being particularly preferred. The peptides may be digests of naturally occurring proteins as is outlined above, random peptides, or "biased" random peptides. By "randomized" or grammatical equivalents herein is meant that each nucleic acid and peptide consists of essentially random nucleotides and amino acids, respectively. Since generally these random peptides (or nucleic acids, discussed below) are chemically synthesized, they may incorporate any nucleotide or amino acid at any position. The synthetic process can be designed to generate randomized proteins or nucleic acids, to allow the formation of all or most of the possible combinations over the length of the sequence, thus forming a library of randomized candidate bioactive proteinaceous agents.

In one embodiment, the library is fully randomized, with no sequence preferences or constants at any position. In a preferred embodiment, the library is biased. That is, some positions within the sequence are either held constant, or are selected from a limited number of possibilities. For example, in a preferred embodiment, the nucleotides or amino acid residues are randomized within a defined class, e.g., of hydrophobic amino acids, hydrophilic residues, sterically biased (either small or large) residues, towards the creation of nucleic acid binding domains, the creation of cysteines, for cross-linking, prolines for SH-3 domains, serines, threonines, tyrosines or histidines for phosphorylation sites, etc., or to purines, etc.

Modulators of prostate cancer can also be nucleic acids, as defined below. As described above generally for proteins, nucleic acid modulating agents may be naturally occurring nucleic acids, random nucleic acids, or "biased" random nucleic acids. For

example, digests of procaryotic or eucaryotic genomes may be used as is outlined above for proteins.

In certain embodiments, the activity of a prostate cancer-associated protein is down-regulated, or entirely inhibited, by the use of antisense polynucleotide, i.e., a nucleic acid complementary to, and which can preferably hybridize specifically to, a coding mRNA nucleic acid sequence, e.g., a prostate cancer protein mRNA, or a subsequence thereof. Binding of the antisense polynucleotide to the mRNA reduces the translation and/or stability of the mRNA.

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In the context of this invention, antisense polynucleotides can comprise naturally-occurring nucleotides, or synthetic species formed from naturally-occurring subunits or their close homologs. Antisense polynucleotides may also have altered sugar moieties or inter-sugar linkages. Exemplary among these are the phosphorothicate and other sulfur containing species which are known for use in the art. Analogs are comprehended by this invention so long as they function effectively to hybridize with the prostate cancer protein mRNA. See, e.g., Isis Pharmaceuticals, Carlsbad, CA; Sequitor, Inc., Natick, MA.

Such antisense polynucleotides can readily be synthesized using recombinant means, or can be synthesized in vitro. Equipment for such synthesis is sold by several vendors, including Applied Biosystems. The preparation of other oligonucleotides such as phosphorothioates and alkylated derivatives is also well known to those of skill in the art.

Antisense molecules as used herein include antisense or sense oligonucleotides. Sense oligonucleotides can, e.g., be employed to block transcription by binding to the anti-sense strand. The antisense and sense oligonucleotide comprise a single-stranded nucleic acid sequence (either RNA or DNA) capable of binding to target mRNA (sense) or DNA (antisense) sequences for prostate cancer molecules. Antisense or sense oligonucleotides, according to the present invention, comprise a fragment generally at least about 14 nucleotides, preferably from about 14 to 30 nucleotides. The ability to derive an antisense or a sense oligonucleotide, based upon a cDNA sequence encoding a given protein is described in, e.g., Stein & Cohen (Cancer Res. 48:2659 (1988 and van der Krol et al. (BioTechniques 6:958 (1988)).

In addition to antisense polynucleotides, ribozymes can be used to target and inhibit transcription of prostate cancer-associated nucleotide sequences. A ribozyme is an

RNA molecule that catalytically cleaves other RNA molecules. Different kinds of ribozymes have been described, including group I ribozymes, hammerhead ribozymes, hairpin ribozymes, RNase P, and axhead ribozymes (see, e.g., Castanotto et al., Adv. in Pharmacology 25: 289-317 (1994) for a general review of the properties of different ribozymes).

The general features of hairpin ribozymes are described, e.g., in Hampel et al., Nucl. Acids Res. 18:299-304 (1990); European Patent Publication No. 0 360 257; U.S. Patent No. 5,254,678. Methods of preparing are well known to those of skill in the art (see, e.g., WO 94/26877; Ojwang et al., Proc. Natl. Acad. Sci. USA 90:6340-6344 (1993); Yamada et al., Human Gene Therapy 1:39-45 (1994); Leavitt et al., Proc. Natl. Acad. Sci. USA 92:699-703 (1995); Leavitt et al., Human Gene Therapy 5:1151-120 (1994); and Yamada et al., Virology 205: 121-126 (1994)).

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Polynucleotide modulators of prostate cancer may be introduced into a cell containing the target nucleotide sequence by formation of a conjugate with a ligand binding molecule, as described in WO 91/04753. Suitable ligand binding molecules include, but are not limited to, cell surface receptors, growth factors, other cytokines, or other ligands that bind to cell surface receptors. Preferably, conjugation of the ligand binding molecule does not substantially interfere with the ability of the ligand binding molecule to bind to its corresponding molecule or receptor, or block entry of the sense or antisense oligonucleotide or its conjugated version into the cell. Alternatively, a polynucleotide modulator of prostate cancer may be introduced into a cell containing the target nucleic acid sequence, e.g., by formation of an polynucleotide-lipid complex, as described in WO 90/10448. It is understood that the use of antisense molecules or knock out and knock in models may also be used in screening assays as discussed above, in addition to methods of treatment.

As noted above, gene expression monitoring is conveniently used to test candidate modultors (e.g., protein, nucleic acid or small molecule). After the candidate agent has been added and the cells allowed to incubate for some period of time, the sample containing a target sequence to be analyzed is added to the biochip. If required, the target sequence is prepared using known techniques. For example, the sample may be treated to lyse the cells, using known lysis buffers, electroporation, etc., with purification and/or amplification such as PCR performed as appropriate. For example, an in vitro transcription

with labels covalently attached to the nucleotides is performed. Generally, the nucleic acids are labeled with biotin-FITC or PE, or with cy3 or cy5.

In a preferred embodiment, the target sequence is labeled with, e.g., a fluorescent, a chemiluminescent, a chemical, or a radioactive signal, to provide a means of detecting the target sequence's specific binding to a probe. The label also can be an enzyme, 5 such as, alkaline phosphatase or horseradish peroxidase, which when provided with an appropriate substrate produces a product that can be detected. Alternatively, the label can be a labeled compound or small molecule, such as an enzyme inhibitor, that binds but is not catalyzed or altered by the enzyme. The label also can be a moiety or compound, such as, an epitope tag or biotin which specifically binds to streptavidin. For the example of biotin, the streptavidin is labeled as described above, thereby, providing a detectable signal for the bound target sequence. Unbound labeled streptavidin is typically removed prior to analysis.

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As will be appreciated by those in the art, these assays can be direct hybridization assays or can comprise "sandwich assays", which include the use of multiple 15 probes, as is generally outlined in U.S. Patent Nos. 5,681,702, 5,597,909, 5,545,730, 5,594,117, 5,591,584, 5,571,670, 5,580,731, 5,571,670, 5,591,584, 5,624,802, 5,635,352, 5,594,118, 5,359,100, 5,124,246 and 5,681,697, all of which are hereby incorporated by reference. In this embodiment, in general, the target nucleic acid is prepared as outlined above, and then added to the biochip comprising a plurality of nucleic acid probes, under conditions that allow the formation of a hybridization complex.

A variety of hybridization conditions may be used in the present invention, including high, moderate and low stringency conditions as outlined above. The assays are generally run under stringency conditions which allows formation of the label probe hybridization complex only in the presence of target. Stringency can be controlled by altering a step parameter that is a thermodynamic variable, including, but not limited to, temperature, formamide concentration, salt concentration, chaotropic salt concentration pH, organic solvent concentration, etc.

These parameters may also be used to control non-specific binding, as is generally outlined in U.S. Patent No. 5,681,697. Thus it may be desirable to perform certain steps at higher stringency conditions to reduce non-specific binding. 30

The reactions outlined herein may be accomplished in a variety of ways.

Components of the reaction may be added simultaneously, or sequentially, in different orders, with preferred embodiments outlined below. In addition, the reaction may include a variety of other reagents. These include salts, buffers, neutral proteins, e.g. albumin, detergents, etc. which may be used to facilitate optimal hybridization and detection, and/or reduce non-specific or background interactions. Reagents that otherwise improve the efficiency of the assay, such as protease inhibitors, nuclease inhibitors, anti-microbial agents, etc., may also be used as appropriate, depending on the sample preparation methods and purity of the target.

The assay data are analyzed to determine the expression levels, and changes in expression levels as between states, of individual genes, forming a gene expression profile.

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Screens are performed to identify modulators of the prostate cancer phenotype. In one embodiment, screening is performed to identify modulators that can induce or suppress a particular expression profile, thus preferably generating the associated phenotype. In another embodiment, e.g., for diagnostic applications, having identified differentially expressed genes important in a particular state, screens can be performed to identify modulators that alter expression of individual genes. In an another embodiment, screening is performed to identify modulators that alter a biological function of the expression product of a differentially expressed gene. Again, having identified the importance of a gene in a particular state, screens are performed to identify agents that bind and/or modulate the biological activity of the gene product.

In addition screens can be done for genes that are induced in response to a candidate agent. After identifying a modulator based upon its ability to suppress a prostate cancer expression pattern leading to a normal expression pattern, or to modulate a single prostate cancer gene expression profile so as to mimic the expression of the gene from normal tissue, a screen as described above can be performed to identify genes that are specifically modulated in response to the agent. Comparing expression profiles between normal tissue and agent treated prostate cancer tissue reveals genes that are not expressed in normal tissue or prostate cancer tissue, but are expressed in agent treated tissue. These agent-specific sequences can be identified and used by methods described herein for prostate cancer genes or proteins. In particular these sequences and the proteins they encode find use in marking or identifying agent treated cells. In addition, antibodies can be raised against the

agent induced proteins and used to target novel therapeutics to the treated prostate cancer tissue sample.

Thus, in one embodiment, a test compound is administered to a population of prostate cancer cells, that have an associated prostate cancer expression profile. By "administration" or "contacting" herein is meant that the candidate agent is added to the cells 5 in such a manner as to allow the agent to act upon the cell, whether by uptake and intracellular action, or by action at the cell surface. In some embodiments, nucleic acid encoding a proteinaceous candidate agent (i.e., a peptide) may be put into a viral construct such as an adenoviral or retroviral construct, and added to the cell, such that expression of the peptide agent is accomplished, e.g., PCT US97/01019. Regulatable gene therapy systems can also be used.

Once the test compound has been administered to the cells, the cells can be washed if desired and are allowed to incubate under preferably physiological conditions for some period of time. The cells are then harvested and a new gene expression profile is generated, as outlined herein.

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Thus, e.g., prostate cancer tissue may be screened for agents that modulate, e.g., induce or suppress the prostate cancer phenotype. A change in at least one gene, preferably many, of the expression profile indicates that the agent has an effect on prostate cancer activity. By defining such a signature for the prostate cancer phenotype, screens for new drugs that alter the phenotype can be devised. With this approach, the drug target need not be known and need not be represented in the original expression screening platform, nor does the level of transcript for the target protein need to change.

In a preferred embodiment, as outlined above, screens may be done on individual genes and gene products (proteins). That is, having identified a particular 25 differentially expressed gene as important in a particular state, screening of modulators of either the expression of the gene or the gene product itself can be done. The gene products of differentially expressed genes are sometimes referred to herein as "prostate cancer proteins" or a "prostate cancer modulatory protein". The prostate cancer modulatory protein may be a fragment, or alternatively, be the full length protein to the fragment encoded by the nucleic acids of Tables 1-16. Preferably, the prostate cancer modulatory protein is a fragment. In a preferred embodiment, the prostate cancer amino acid sequence which is used to determine

sequence identity or similarity is encoded by a nucleic acid of Tables 1-16. In another embodiment, the sequences are naturally occurring allelic variants of a protein encoded by a nucleic acid of Tables 1-16. In another embodiment, the sequences are sequence variants as further described herein.

Preferably, the prostate cancer modulatory protein is a fragment of approximately 14 to 24 amino acids long. More preferably the fragment is a soluble fragment. Preferably, the fragment includes a non-transmembrane region. In a preferred embodiment, the fragment has an N-terminal Cys to aid in solubility. In one embodiment, the C-terminus of the fragment is kept as a free acid and the N-terminus is a free amine to aid in coupling, i.e., to cysteine.

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In one embodiment the prostate cancer proteins are conjugated to an immunogenic agent as discussed herein. In one embodiment the prostate cancer protein is conjugated to BSA.

Measurements of prostate cancer polypeptide activity, or of prostate cancer or the prostate cancer phenotype can be performed using a variety of assays. For example, the effects of the test compounds upon the function of the prostate cancer polypeptides can be measured by examining parameters described above. A suitable physiological change that affects activity can be used to assess the influence of a test compound on the polypeptides of this invention. When the functional consequences are determined using intact cells or animals, one can also measure a variety of effects such as, in the case of prostate cancer associated with tumors, tumor growth, tumor metastasis, neovascularization, hormone release, transcriptional changes to both known and uncharacterized genetic markers (e.g., northern blots), changes in cell metabolism such as cell growth or pH changes, and changes in intracellular second messengers such as cGMP. In the assays of the invention, mammalian prostate cancer polypeptide is typically used, e.g., mouse, preferably human.

Assays to identify compounds with modulating activity can be performed in vitro. For example, a prostate cancer polypeptide is first contacted with a potential modulator and incubated for a suitable amount of time, e.g., from 0.5 to 48 hours. In one embodiment, the prostate cancer polypeptide levels are determined in vitro by measuring the level of protein or mRNA. The level of protein is measured using immunoassays such as western blotting, ELISA and the like with an antibody that selectively binds to the prostate cancer

polypeptide or a fragment thereof. For measurement of mRNA, amplification, e.g., using PCR, LCR, or hybridization assays, e.g., northern hybridization, RNAse protection, dot blotting, are preferred. The level of protein or mRNA is detected using directly or indirectly labeled detection agents, e.g., fluorescently or radioactively labeled nucleic acids, radioactively or enzymatically labeled antibodies, and the like, as described herein.

Alternatively, a reporter gene system can be devised using the prostate cancer protein promoter operably linked to a reporter gene such as luciferase, green fluorescent protein, CAT, or  $\beta$ -gal. The reporter construct is typically transfected into a cell. After treatment with a potential modulator, the amount of reporter gene transcription, translation, or activity is measured according to standard techniques known to those of skill in the art.

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In a preferred embodiment, as outlined above, screens may be done on individual genes and gene products (proteins). That is, having identified a particular differentially expressed gene as important in a particular state, screening of modulators of the expression of the gene or the gene product itself can be done. The gene products of differentially expressed genes are sometimes referred to herein as "prostate cancer proteins."

The prostate cancer protein may be a fragment, or alternatively, be the full length protein to a fragment shown herein.

In one embodiment, screening for modulators of expression of specific genes is performed. Typically, the expression of only one or a few genes are evaluated. In another embodiment, screens are designed to first find compounds that bind to differentially expressed proteins. These compounds are then evaluated for the ability to modulate differentially expressed activity. Moreover, once initial candidate compounds are identified, variants can be further screened to better evaluate structure activity relationships.

In a preferred embodiment, binding assays are done. In general, purified or isolated gene product is used; that is, the gene products of one or more differentially expressed nucleic acids are made. For example, antibodies are generated to the protein gene products, and standard immunoassays are run to determine the amount of protein present. Alternatively, cells comprising the prostate cancer proteins can be used in the assays.

Thus, in a preferred embodiment, the methods comprise combining a prostate cancer protein and a candidate compound, and determining the binding of the compound to the prostate cancer protein. Preferred embodiments utilize the human prostate cancer protein,

although other mammalian proteins may also be used, e.g. for the development of animal models of human disease. In some embodiments, as outlined herein, variant or derivative prostate cancer proteins may be used.

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Generally, in a preferred embodiment of the methods herein, the prostate cancer protein or the candidate agent is non-diffusably bound to an insoluble support having isolated sample receiving areas (e.g. a microtiter plate, an array, etc.). The insoluble supports may be made of any composition to which the compositions can be bound, is readily separated from soluble material, and is otherwise compatible with the overall method of screening. The surface of such supports may be solid or porous and of any convenient shape. Examples of suitable insoluble supports include microtiter plates, arrays, membranes and beads. These are typically made of glass, plastic (e.g., polystyrene), polysaccharides, nylon or nitrocellulose, teflon™, etc. Microtiter plates and arrays are especially convenient because a large number of assays can be carried out simultaneously, using small amounts of reagents and samples. The particular manner of binding of the composition is not crucial so long as it is compatible with the reagents and overall methods of the invention, maintains the activity of the composition and is nondiffusable. Preferred methods of binding include the use of antibodies (which do not sterically block either the ligand binding site or activation sequence when the protein is bound to the support), direct binding to "sticky" or ionic supports, chemical crosslinking, the synthesis of the protein or agent on the surface, etc. Following binding of the protein or agent, excess unbound material is removed by washing. The sample receiving areas may then be blocked through incubation with bovine serum albumin (BSA), casein or other innocuous protein or other moiety.

In a preferred embodiment, the prostate cancer protein is bound to the support, and a test compound is added to the assay. Alternatively, the candidate agent is bound to the support and the prostate cancer protein is added. Novel binding agents include specific antibodies, non-natural binding agents identified in screens of chemical libraries, peptide analogs, etc. Of particular interest are screening assays for agents that have a low toxicity for human cells. A wide variety of assays may be used for this purpose, including labeled in vitro protein-protein binding assays, electrophoretic mobility shift assays, immunoassays for protein binding, functional assays (phosphorylation assays, etc.) and the like.

The determination of the binding of the test modulating compound to the prostate cancer protein may be done in a number of ways. In a preferred embodiment, the compound is labeled, and binding determined directly, e.g., by attaching all or a portion of the prostate cancer protein to a solid support, adding a labeled candidate agent (e.g., a fluorescent label), washing off excess reagent, and determining whether the label is present on the solid support. Various blocking and washing steps may be utilized as appropriate.

In some embodiments, only one of the components is labeled, e.g., the proteins (or proteinaceous candidate compounds) can be labeled. Alternatively, more than one component can be labeled with different labels, e.g., <sup>125</sup>I for the proteins and a fluorophor for the compound. Proximity reagents, e.g., quenching or energy transfer reagents are also useful.

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In one embodiment, the binding of the test compound is determined by competitive binding assay. The competitor is a binding moiety known to bind to the target molecule (i.e., a prostate cancer protein), such as an antibody, peptide, binding partner, ligand, etc. Under certain circumstances, there may be competitive binding between the compound and the binding moiety, with the binding moiety displacing the compound. In one embodiment, the test compound is labeled. Either the compound, or the competitor, or both, is added first to the protein for a time sufficient to allow binding, if present. Incubations may be performed at a temperature which facilitates optimal activity, typically between 4 and 40°C. Incubation periods are typically optimized, e.g., to facilitate rapid high throughput screening. Typically between 0.1 and 1 hour will be sufficient. Excess reagent is generally removed or washed away. The second component is then added, and the presence or absence of the labeled component is followed, to indicate binding.

In a preferred embodiment, the competitor is added first, followed by the test compound. Displacement of the competitor is an indication that the test compound is binding to the prostate cancer protein and thus is capable of binding to, and potentially modulating, the activity of the prostate cancer protein. In this embodiment, either component can be labeled. Thus, e.g., if the competitor is labeled, the presence of label in the wash solution indicates displacement by the agent. Alternatively, if the test compound is labeled, the presence of the label on the support indicates displacement.

In an alternative embodiment, the test compound is added first, with incubation and washing, followed by the competitor. The absence of binding by the competitor may indicate that the test compound is bound to the prostate cancer protein with a higher affinity. Thus, if the test compound is labeled, the presence of the label on the support, coupled with a lack of competitor binding, may indicate that the test compound is capable of binding to the prostate cancer protein.

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In a preferred embodiment, the methods comprise differential screening to identity agents that are capable of modulating the activity of the prostate cancer proteins. In this embodiment, the methods comprise combining a prostate cancer protein and a competitor in a first sample. A second sample comprises a test compound, a prostate cancer protein, and a competitor. The binding of the competitor is determined for both samples, and a change, or difference in binding between the two samples indicates the presence of an agent capable of binding to the prostate cancer protein and potentially modulating its activity. That is, if the binding of the competitor is different in the second sample relative to the first sample, the agent is capable of binding to the prostate cancer protein.

Alternatively, differential screening is used to identify drug candidates that bind to the native prostate cancer protein, but cannot bind to modified prostate cancer proteins. The structure of the prostate cancer protein may be modeled, and used in rational drug design to synthesize agents that interact with that site. Drug candidates that affect the activity of a prostate cancer protein are also identified by screening drugs for the ability to either enhance or reduce the activity of the protein.

Positive controls and negative controls may be used in the assays. Preferably control and test samples are performed in at least triplicate to obtain statistically significant results. Incubation of all samples is for a time sufficient for the binding of the agent to the protein. Following incubation, samples are washed free of non-specifically bound material and the amount of bound, generally labeled agent determined. For example, where a radiolabel is employed, the samples may be counted in a scintillation counter to determine the amount of bound compound.

A variety of other reagents may be included in the screening assays. These include reagents like salts, neutral proteins, e.g. albumin, detergents, etc. which may be used to facilitate optimal protein-protein binding and/or reduce non-specific or background

interactions. Also reagents that otherwise improve the efficiency of the assay, such as protease inhibitors, nuclease inhibitors, anti-microbial agents, etc., may be used. The mixture of components may be added in an order that provides for the requisite binding.

In a preferred embodiment, the invention provides methods for screening for a compound capable of modulating the activity of a prostate cancer protein. The methods comprise adding a test compound, as defined above, to a cell comprising prostate cancer proteins. Preferred cell types include almost any cell. The cells contain a recombinant nucleic acid that encodes a prostate cancer protein. In a preferred embodiment, a library of candidate agents are tested on a plurality of cells.

In one aspect, the assays are evaluated in the presence or absence or previous or subsequent exposure of physiological signals, e.g. hormones, antibodies, peptides, antigens, cytokines, growth factors, action potentials, pharmacological agents including chemotherapeutics, radiation, carcinogenics, or other cells (i.e. cell-cell contacts). In another example, the determinations are determined at different stages of the cell cycle process.

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In this way, compounds that modulate prostate cancer agents are identified.

Compounds with pharmacological activity are able to enhance or interfere with the activity of the prostate cancer protein. Once identified, similar structures are evaluated to identify critical structural feature of the compound.

In one embodiment, a method of inhibiting prostate cancer cell division is provided. The method comprises administration of a prostate cancer inhibitor. In another embodiment, a method of inhibiting prostate cancer is provided. The method comprises administration of a prostate cancer inhibitor. In a further embodiment, methods of treating cells or individuals with prostate cancer are provided. The method comprises administration of a prostate cancer inhibitor.

In one embodiment, a prostate cancer inhibitor is an antibody as discussed above. In another embodiment, the prostate cancer inhibitor is an antisense molecule.

A variety of cell growth, proliferation, and metastasis assays are known to those of skill in the art, as described below.

Soft agar growth or colony formation in suspension

Normal cells require a solid substrate to attach and grow. When the cells are transformed, they lose this phenotype and grow detached from the substrate. For example,

transformed cells can grow in stirred suspension culture or suspended in semi-solid media, such as semi-solid or soft agar. The transformed cells, when transfected with tumor suppressor genes, regenerate normal phenotype and require a solid substrate to attach and grow. Soft agar growth or colony formation in suspension assays can be used to identify modulators of prostate cancer sequences, which when expressed in host cells, inhibit abnormal cellular proliferation and transformation. A therapeutic compound would reduce or eliminate the host cells' ability to grow in stirred suspension culture or suspended in semi-solid media, such as semi-solid or soft.

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Techniques for soft agar growth or colony formation in suspension assays are described in Freshney, Culture of Animal Cells a Manual of Basic Technique (3<sup>rd</sup> ed., 1994), herein incorporated by reference. See also, the methods section of Garkavtsev et al. (1996), supra. herein incorporated by reference.

Contact inhibition and density limitation of growth

Normal cells typically grow in a flat and organized pattern in a petri dish until

they touch other cells. When the cells touch one another, they are contact inhibited and stop growing. When cells are transformed, however, the cells are not contact inhibited and continue to grow to high densities in disorganized foci. Thus, the transformed cells grow to a higher saturation density than normal cells. This can be detected morphologically by the formation of a disoriented monolayer of cells or rounded cells in foci within the regular pattern of normal surrounding cells. Alternatively, labeling index with (<sup>3</sup>H)-thymidine at saturation density can be used to measure density limitation of growth. See Freshney (1994), supra. The transformed cells, when transfected with tumor suppressor genes, regenerate a normal phenotype and become contact inhibited and would grow to a lower density.

In this assay, labeling index with ( $^3$ H)-thymidine at saturation density is a preferred method of measuring density limitation of growth. Transformed host cells are transfected with a prostate cancer-associated sequence and are grown for 24 hours at saturation density in non-limiting medium conditions. The percentage of cells labeling with ( $^3$ H)-thymidine is determined autoradiographically. See, Freshney (1994), supra.

### Growth factor or serum dependence

Transformed cells have a lower serum dependence than their normal counterparts (see, e.g., Temin, J. Natl. Cancer Insti. 37:167-175 (1966); Eagle et al., J. Exp. Med. 131:836-879 (1970)); Freshney, supra. This is in part due to release of various growth factors by the transformed cells. Growth factor or serum dependence of transformed host cells can be compared with that of control.

### Tumor specific markers levels

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Tumor cells release an increased amount of certain factors (hereinafter "tumor specific markers") than their normal counterparts. For example, plasminogen activator (PA) is released from human glioma at a higher level than from normal brain cells (see, e.g., Gullino, Angiogenesis, tumor vascularization, and potential interference with tumor growth. in Biological Responses in Cancer, pp. 178-184 (Mihich (ed.) 1985)). Similarly, Tumor angiogenesis factor (TAF) is released at a higher level in tumor cells than their normal counterparts. See, e.g., Folkman, Angiogenesis and Cancer, Sem Cancer Biol. (1992)).

Various techniques which measure the release of these factors are described in Freshney (1994), supra. Also, see, Unkless et al., J. Biol. Chem. 249:4295-4305 (1974); Strickland & Beers, J. Biol. Chem. 251:5694-5702 (1976); Whur et al., Br. J. Cancer 42:305-312 (1980); Gullino, Anglogenesis, tumor vascularization, and potential interference with tumor growth. in Biological Responses in Cancer, pp. 178-184 (Mihich (ed.) 1985); Freshney Anticancer Res. 5:111-130 (1985).

## Invasiveness into Matrigel

The degree of invasiveness into Matrigel or some other extracellular matrix constituent can be used as an assay to identify compounds that modulate prostate cancer-associated sequences. Tumor cells exhibit a good correlation between malignancy and invasiveness of cells into Matrigel or some other extracellular matrix constituent. In this assay, tumorigenic cells are typically used as host cells. Expression of a tumor suppressor gene in these host cells would decrease invasiveness of the host cells.

Techniques described in Freshney (1994), *supra*, can be used. Briefly, the level of invasion of host cells can be measured by using filters coated with Matrigel or some

other extracellular matrix constituent. Penetration into the gel, or through to the distal side of the filter, is rated as invasiveness, and rated histologically by number of cells and distance moved, or by prelabeling the cells with <sup>125</sup>I and counting the radioactivity on the distal side of the filter or bottom of the dish. See, e.g., Freshney (1984), supra.

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### Tumor growth in vivo

Effects of prostate cancer-associated sequences on cell growth can be tested in transgenic or immune-suppressed mice. Knock-out transgenic mice can be made, in which the prostate cancer gene is disrupted or in which a prostate cancer gene is inserted. Knock-out transgenic mice can be made by insertion of a marker gene or other heterologous gene into the endogenous prostate cancer gene site in the mouse genome via homologous recombination. Such mice can also be made by substituting the endogenous prostate cancer gene with a mutated version of the prostate cancer gene, or by mutating the endogenous prostate cancer gene, e.g., by exposure to carcinogens.

A DNA construct is introduced into the nuclei of embryonic stem cells. Cells containing the newly engineered genetic lesion are injected into a host mouse embryo, which is re-implanted into a recipient female. Some of these embryos develop into chimeric mice that possess germ cells partially derived from the mutant cell line. Therefore, by breeding the chimeric mice it is possible to obtain a new line of mice containing the introduced genetic lesion (see, e.g., Capecchi et al., Science 244:1288 (1989)). Chimeric targeted mice can be derived according to Hogan et al., Manipulating the Mouse Embryo: A Laboratory Manual, Cold Spring Harbor Laboratory (1988) and Teratocarcinomas and Embryonic Stem Cells: A Practical Approach, Robertson, ed., IRL Press, Washington, D.C., (1987).

Alternatively, various immune-suppressed or immune-deficient host animals can be used. For example, genetically athymic "nude" mouse (see, e.g., Giovanella et al., J. Nail. Cancer Inst. 52:921 (1974)), a SCID mouse, a thymectomized mouse, or an irradiated mouse (see, e.g., Bradley et al., Br. J. Cancer 38:263 (1978); Selby et al., Br. J. Cancer 41:52 (1980)) can be used as a host. Transplantable tumor cells (typically about 10<sup>6</sup> cells) injected into isogenic hosts will produce invasive tumors in a high proportions of cases, while normal cells of similar origin will not. In hosts which developed invasive tumors, cells expressing a prostate cancer-associated sequences are injected subcutaneously. After a

suitable length of time, preferably 4-8 weeks, tumor growth is measured (e.g., by volume or by its two largest dimensions) and compared to the control. Tumors that have statistically significant reduction (using, e.g., Student's T test) are said to have inhibited growth.

### 5 Methods of identifying variant prostate cancer-associated sequences

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Without being bound by theory, expression of various prostate cancer sequences is correlated with prostate cancer. Accordingly, disorders based on mutant or variant prostate cancer genes may be determined. In one embodiment, the invention provides methods for identifying cells containing variant prostate cancer genes, e.g., determining all or part of the sequence of at least one endogenous prostate cancer genes in a cell. This may be accomplished using any number of sequencing techniques. In a preferred embodiment, the invention provides methods of identifying the prostate cancer genotype of an individual, e.g., determining all or part of the sequence of at least one prostate cancer gene of the individual. This is generally done in at least one tissue of the individual, and may include the evaluation of a number of tissues or different samples of the same tissue. The method may include comparing the sequence of the sequenced prostate cancer gene to a known prostate cancer gene. i.e., a wild-type gene.

The sequence of all or part of the prostate cancer gene can then be compared to the sequence of a known prostate cancer gene to determine if any differences exist. This can be done using any number of known homology programs, such as Bestfit, etc. In a preferred embodiment, the presence of a difference in the sequence between the prostate cancer gene of the patient and the known prostate cancer gene correlates with a disease state or a propensity for a disease state, as outlined herein.

In a preferred embodiment, the prostate cancer genes are used as probes to determine the number of copies of the prostate cancer gene in the genome.

In another preferred embodiment, the prostate cancer genes are used as probes to determine the chromosomal localization of the prostate cancer genes. Information such as chromosomal localization finds use in providing a diagnosis or prognosis in particular when chromosomal abnormalities such as translocations, and the like are identified in the prostate cancer gene locus.

### Administration of pharmaceutical and vaccine compositions

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In one embodiment, a therapeutically effective dose of a prostate cancer protein or modulator thereof, is administered to a patient. By "therapeutically effective dose" herein is meant a dose that produces effects for which it is administered. The exact dose will depend on the purpose of the treatment, and will be ascertainable by one skilled in the art 5 using known techniques (e.g., Ansel et al., Pharmaceutical Dosage Forms and Drug Delivery: Lieberman, Pharmaceutical Dosage Forms (vols. 1-3, 1992), Dekker, ISBN 0824770846, 082476918X, 0824712692, 0824716981; Lloyd, The Art, Science and Technology of Pharmaceutical Compounding (1999); and Pickar, Dosage Calculations (1999)). As is known in the art, adjustments for prostate cancer degradation, systemic versus localized delivery, and rate of new protease synthesis, as well as the age, body weight, general health, sex, diet, time of administration, drug interaction and the severity of the condition may be necessary, and will be ascertainable with routine experimentation by those skilled in the art, U.S. Patent Application N. 09/687,576, further discloses the use of 15 compositions and methods of diagnosis and treatment in prostate cancer is hereby expressly incorporated by reference.

A "patient" for the purposes of the present invention includes both humans and other animals, particularly mammals. Thus the methods are applicable to both human therapy and veterinary applications. In the preferred embodiment the patient is a mammal, preferably a primate, and in the most preferred embodiment the patient is human.

The administration of the prostate cancer proteins and modulators thereof of the present invention can be done in a variety of ways as discussed above, including, but not limited to, orally, subcutaneously, intravenously, intranasally, transdermally, intraperitoneally, intramuscularly, intrapulmonary, vaginally, rectally, or intraocularly. In some instances, e.g., in the treatment of wounds and inflammation, the prostate cancer proteins and modulators may be directly applied as a solution or spray.

The pharmaceutical compositions of the present invention comprise a prostate cancer protein in a form suitable for administration to a patient. In the preferred embodiment, the pharmaceutical compositions are in a water soluble form, such as being present as pharmaceutically acceptable salts, which is meant to include both acid and base addition salts. "Pharmaceutically acceptable acid addition salt" refers to those salts that retain the

biological effectiveness of the free bases and that are not biologically or otherwise undesirable, formed with inorganic acids such as hydrochloric acid, hydrobromic acid, sulfuric acid, nitric acid, phosphoric acid and the like, and organic acids such as acetic acid, propionic acid, glycolic acid, pyruvic acid, oxalic acid, maleic acid, malonic acid, succinic acid, fumaric acid, tentanesulfonic acid, ethanesulfonic acid, ethanesulfonic acid, ethanesulfonic acid, salicylic acid and the like. "Pharmaceutically acceptable base addition salts" include those derived from inorganic bases such as sodium, potassium, lithium, ammonium, calcium, magnesium, iron, zinc, copper, manganese, aluminum salts and the like. Particularly preferred are the ammonium, potassium, sodium, calcium, and magnesium salts. Salts derived from pharmaceutically acceptable organic non-toxic bases include salts of primary, secondary, and tertiary amines, substituted amines including naturally occurring substituted amines, cyclic amines and basic ion exchange resins, such as isopropylamine, trimethylamine, diethylamine, triethylamine, tripropylamine, and ethanolamine.

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The pharmaceutical compositions may also include one or more of the following: carrier proteins such as serum albumin; buffers; fillers such as microcrystalline cellulose, lactose, corn and other starches; binding agents; sweeteners and other flavoring agents; coloring agents; and polyethylene glycol.

The pharmaceutical compositions can be administered in a variety of unit dosage forms depending upon the method of administration. For example, unit dosage forms suitable for oral administration include, but are not limited to, powder, tablets, pills, capsules and lozenges. It is recognized that prostate cancer protein modulators (e.g., antibodies, antisense constructs, ribozymes, small organic molecules, etc.) when administered orally, should be protected from digestion. This is typically accomplished either by complexing the molecule(s) with a composition to render it resistant to acidic and enzymatic hydrolysis, or by packaging the molecule(s) in an appropriately resistant carrier, such as a liposome or a protection barrier. Means of protecting agents from digestion are well known in the art.

The compositions for administration will commonly comprise a prostate cancer protein modulator dissolved in a pharmaceutically acceptable carrier, preferably an aqueous carrier. A variety of aqueous carriers can be used, e.g., buffered saline and the like. These solutions are sterile and generally free of undesirable matter. These compositions may

be sterilized by conventional, well known sterilization techniques. The compositions may contain pharmaceutically acceptable auxiliary substances as required to approximate physiological conditions such as pH adjusting and buffering agents, toxicity adjusting agents and the like, e.g., sodium acetate, sodium chloride, potassium chloride, calcium chloride, sodium lactate and the like. The concentration of active agent in these formulations can vary widely, and will be selected primarily based on fluid volumes, viscosities, body weight and the like in accordance with the particular mode of administration selected and the patient's needs (e.g., Remington's Pharmaceutical Science (15th ed., 1980) and Goodman & Gillman, The Pharmacologial Basis of Therapeutics (Hardman et al., eds., 1996)).

Thus, a typical pharmaceutical composition for intraveneous administration would be about 0.1 to 10 mg per patient per day. Dosages from 0.1 up to about 100 mg per patient per day may be used, particularly when the drug is administered to a secluded site and not into the blood stream, such as into a body cavity or into a lumen of an organ. Substantially higher dosages are possible in topical administration. Actual methods for preparing parenterally administrable compositions will be known or apparent to those skilled in the art, e.g., Remington's Pharmaceutical Science and Goodman and Gillman, The Pharmacologial Basis of Therapeutics, supra.

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The compositions containing modulators of prostate cancer proteins can be administered for therapeutic or prophylactic treatments. In therapeutic applications, compositions are administered to a patient suffering from a disease (e.g., a cancer) in an amount sufficient to cure or at least partially arrest the disease and its complications. An amount adequate to accomplish this is defined as a "therapeutically effective dose." Amounts effective for this use will depend upon the severity of the disease and the general state of the patient's health. Single or multiple administrations of the compositions may be administered depending on the dosage and frequency as required and tolerated by the patient. In any event, the composition should provide a sufficient quantity of the agents of this invention to effectively treat the patient. An amount of modulator that is capable of preventing or slowing the development of cancer in a mammal is referred to as a "prophylactically effective dose."

The particular dose required for a prophylactic treatment will depend upon the medical condition and history of the mammal, the particular cancer being prevented, as well as other factors such as age, weight, gender, administration route, efficiency, etc. Such prophylactic

treatments may be used, e.g., in a mammal who has previously had cancer to prevent a recurrence of the cancer, or in a mammal who is suspected of having a significant likelihood of developing cancer.

It will be appreciated that the present prostate cancer protein-modulating compounds can be administered alone or in combination with additional prostate cancer modulating compounds or with other therapeutic agent, e.g., other anti-cancer agents or treatments.

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In numerous embodiments, one or more nucleic acids, e.g., polynucleotides comprising nucleic acid sequences set forth in Tables 1-16, such as antisense polynucleotides or ribozymes, will be introduced into cells, in vitro or in vivo. The present invention provides methods, reagents, vectors, and cells useful for expression of prostate cancer-associated polypeptides and nucleic acids using in vitro (cell-free), ex vivo or in vivo (cell or organism-based) recombinant expression systems.

The particular procedure used to introduce the nucleic acids into a host cell for expression of a protein or nucleic acid is application specific. Many procedures for introducing foreign nucleotide sequences into host cells may be used. These include the use of calcium phosphate transfection, spheroplasts, electroporation, liposomes, microinjection, plasma vectors, viral vectors and any of the other well known methods for introducing cloned genomic DNA, cDNA, synthetic DNA or other foreign genetic material into a host cell (see, e.g., Berger & Kimmel, Guide to Molecular Cloning Techniques, Methods in Enzymology volume 152 (Berger), Ausubel et al., eds., Current Protocols (supplemented through 1999), and Sambrook et al., Molecular Cloning - A Laboratory Manual (2nd ed., Vol. 1-3, 1989).

In a preferred embodiment, prostate cancer proteins and modulators are administered as therapeutic agents, and can be formulated as outlined above. Similarly, prostate cancer genes (including both the full-length sequence, partial sequences, or regulatory sequences of the prostate cancer coding regions) can be administered in a gene therapy application. These prostate cancer genes can include antisense applications, either as gene therapy (i.e. for incorporation into the genome) or as antisense compositions, as will be appreciated by those in the art.

Prostate cancer polypeptides and polynucleotides can also be administered as vaccine compositions to stimulate HTL, CTL and antibody responses.. Such vaccine

compositions can include, e.g., lipidated peptides (see, e.g., Vitiello, A. et al., J. Clin. Invest. 95:341 (1995)), pentide compositions encapsulated in poly(DL-lactide-co-glycolide) ("PLG") microspheres (see, e.g., Eldridge, et al., Molec. Immunol. 28:287-294, (1991); Alonso et al., Vaccine 12:299-306 (1994); Jones et al., Vaccine 13:675-681 (1995)), peptide compositions 5 contained in immune stimulating complexes (ISCOMS) (see, e.g., Takahashi et al., Nature 344:873-875 (1990); Hu et al., Clin Exp Immunol. 113:235-243 (1998)), multiple antigen peptide systems (MAPs) (see, e.g., Tam, Proc. Natl. Acad. Sci. U.S.A. 85:5409-5413 (1988); Tam. J. Immunol. Methods 196:17-32 (1996)), peptides formulated as multivalent peptides; peptides for use in ballistic delivery systems, typically crystallized peptides, viral delivery vectors (Perkus, et al., In: Concepts in vaccine development (Kaufmann, ed., p. 379, 1996); 10 Chakrabarti, et al., Nature 320:535 (1986); Hu et al., Nature 320:537 (1986); Kieny, et al., AIDS Bio/Technology 4:790 (1986); Top et al., J. Infect. Dis. 124:148 (1971); Chanda et al., Virology 175:535 (1990)), particles of viral or synthetic origin (see, e.g., Kofler et al., J. Immunol, Methods, 192:25 (1996); Eldridge et al., Sem. Hematol, 30:16 (1993); Falo et al., 15 Nature Med. 7:649 (1995)), adjuvants (Warren et al., Annu. Rev. Immunol. 4:369 (1986); Gupta et al., Vaccine 11:293 (1993)), liposomes (Reddy et al., J. Immunol. 148:1585 (1992); Rock, Immunol. Today 17:131 (1996)), or, naked or particle absorbed cDNA (Ulmer, et al., Science 259:1745 (1993); Robinson et al., Vaccine 11:957 (1993); Shiver et al., In: Concepts in vaccine development (Kaufmann, ed., p. 423, 1996); Cease & Berzofsky, Annu, Rev. Immunol. 12:923 (1994) and Eldridge et al., Sem. Hematol. 30:16 (1993)). Toxin-targeted 20 delivery technologies, also known as receptor mediated targeting, such as those of Avant Immunotherapeutics, Inc. (Needham, Massachusetts) may also be used.

Vaccine compositions often include adjuvants. Many adjuvants contain a substance designed to protect the antigen from rapid catabolism, such as aluminum hydroxide or mineral oil, and a stimulator of immune responses, such as lipid A, Bortadella pertussis or Mycobacterium tuberculosis derived proteins. Certain adjuvants are commercially available as, e.g., Freund's Incomplete Adjuvant and Complete Adjuvant (Difco Laboratories, Detroit, MI); Merck Adjuvant 65 (Merck and Company, Inc., Rahway, NJ); AS-2 (SmithKline Beecham, Philadelphia, PA); aluminum salts such as aluminum hydroxide gel (alum) or aluminum phosphate; salts of calcium, iron or zinc; an insoluble suspension of acylated tyrosine; acylated sugars; cationically or anionically derivatized polysaccharides;

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polyphosphazenes; biodegradable microspheres; monophosphoryl lipid A and quil A. Cytokines, such as GM-CSF, interleukin-2, -7, -12, and other like growth factors, may also be used as adjuvants.

Vaccines can be administered as nucleic acid compositions wherein DNA or RNA encoding one or more of the polypeptides, or a fragment thereof, is administered to a patient. This approach is described, for instance, in Wolff et. al., Science 247:1465 (1990) as well as U.S. Patent Nos. 5,580,859; 5,589,466; 5,804,566; 5,739,118; 5,736,524; 5,679,647; WO 98/04720; and in more detail below. Examples of DNA-based delivery technologies include "naked DNA", facilitated (bupivicaine, polymers, peptide-mediated) delivery, cationic lipid complexes, and particle-mediated ("gene gun") or pressure-mediated delivery 10 (see, e.g., U.S. Patent No. 5,922,687).

For therapeutic or prophylactic immunization purposes, the peptides of the invention can be expressed by viral or bacterial vectors. Examples of expression vectors include attenuated viral hosts, such as vaccinia or fowlpox. This approach involves the use of 15 vaccinia virus, e.g., as a vector to express nucleotide sequences that encode prostate cancer polypeptides or polypeptide fragments. Upon introduction into a host, the recombinant vaccinia virus expresses the immunogenic peptide, and thereby elicits an immune response. Vaccinia vectors and methods useful in immunization protocols are described in, e.g., U.S. Patent No. 4,722,848. Another vector is BCG (Bacille Calmette Guerin). BCG vectors are described in Stover et al., Nature 351:456-460 (1991). A wide variety of other vectors useful for the rapeutic administration or immunization e.g. adeno and adeno-associated virus vectors, retroviral vectors, Salmonella typhi vectors, detoxified anthrax toxin vectors, and the like, will be apparent to those skilled in the art from the description herein (see, e.g., Shata et al., Mol Med Today 6:66-71 (2000); Shedlock et al., J Leukoc Biol 68:793-806 (2000); Hipp et al., In Vivo 14:571-85 (2000)),

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Methods for the use of genes as DNA vaccines are well known, and include placing a prostate cancer gene or portion of a prostate cancer gene under the control of a regulatable promoter or a tissue-specific promoter for expression in a prostate cancer patient. The prostate cancer gene used for DNA vaccines can encode full-length prostate cancer proteins, but more preferably encodes portions of the prostate cancer proteins including peptides derived from the prostate cancer protein. In one embodiment, a patient is

immunized with a DNA vaccine comprising a plurality of nucleotide sequences derived from a prostate cancer gene. For example, prostate cancer-associated genes or sequence encoding subfragments of a prostate cancer protein are introduced into expression vectors and tested for their immunogenicity in the context of Class I MHC and an ability to generate cytotoxic T cell responses. This procedure provides for production of cytotoxic T cell responses against cells which present antigen, including intracellular epitopes.

In a preferred embodiment, the DNA vaccines include a gene encoding an adjuvant molecule with the DNA vaccine. Such adjuvant molecules include cytokines that increase the immunogenic response to the prostate cancer polypeptide encoded by the DNA vaccine. Additional or alternative adjuvants are available.

In another preferred embodiment prostate cancer genes find use in generating animal models of prostate cancer. When the prostate cancer gene identified is repressed or diminished in cancer tissue, gene therapy technology, e.g., wherein antisense RNA directed to the prostate cancer gene will also diminish or repress expression of the gene. Animal models of prostate cancer find use in screening for modulators of a prostate cancer-associated sequence or modulators of prostate cancer. Similarly, transgenic animal technology including gene knockout technology, e.g. as a result of homologous recombination with an appropriate gene targeting vector, will result in the absence or increased expression of the prostate cancer protein. When desired, tissue-specific expression or knockout of the prostate cancer protein may be necessary.

It is also possible that the prostate cancer protein is overexpressed in prostate cancer. As such, transgenic animals can be generated that overexpress the prostate cancer protein. Depending on the desired expression level, promoters of various strengths can be employed to express the transgene. Also, the number of copies of the integrated transgene can be determined and compared for a determination of the expression level of the transgene. Animals generated by such methods find use as animal models of prostate cancer and are additionally useful in screening for modulators to treat prostate cancer.

### Kits for Use in Diagnostic and/or Prognostic Applications

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For use in diagnostic, research, and therapeutic applications suggested above, kits are also provided by the invention. In the diagnostic and research applications such kits

may include any or all of the following: assay reagents, buffers, prostate cancer-specific nucleic acids or antibodies, hybridization probes and/or primers, antisense polynucleotides, ribozymes, dominant negative prostate cancer polypeptides or polynucleotides, small molecules inhibitors of prostate cancer-associated sequences etc. A therapeutic product may include sterile saline or another pharmaceutically acceptable emulsion and suspension base.

In addition, the kits may include instructional materials containing directions (i.e., protocols) for the practice of the methods of this invention. While the instructional materials typically comprise written or printed materials they are not limited to such. Any medium capable of storing such instructions and communicating them to an end user is contemplated by this invention. Such media include, but are not limited to electronic storage media (e.g., magnetic discs, tapes, cartridges, chips), optical media (e.g., CD ROM), and the like. Such media may include addresses to internet sites that provide such instructional materials.

The present invention also provides for kits for screening for modulators of prostate cancer-associated sequences. Such kits can be prepared from readily available materials and reagents. For example, such kits can comprise one or more of the following materials: a prostate cancer-associated polypeptide or polynucleotide, reaction tubes, and instructions for testing prostate cancer-associated activity. Optionally, the kit contains biologically active prostate cancer protein. A wide variety of kits and components can be prepared according to the present invention, depending upon the intended user of the kit and the particular needs of the user. Diagnosis would typically involve evaluation of a plurality of genes or products. The genes will be selected based on correlations with important parameters in disease which may be identified in historical or outcome data.

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#### EXAMPLES

### Example 1: Tissue Preparation, Labeling Chips, and Fingerprints

## Purifying total RNA from tissue sample using TRIzol Reagent

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The sample weight is first estimated. The tissue samples are homogenized in 1 ml of TRIzol per 50 mg of tissue using a homogenizer (e.g., Polytron 3100). The size of the generator/probe used depends upon the sample amount. A generator that is too large for the amount of tissue to be homogenized will cause a loss of sample and lower RNA yield. A larger generator (e.g., 20 mm) is suitable for tissue samples weighing more than 0.6 g. Fill tubes should not be overfilled. If the working volume is greater than 2 ml and no greater than 10 ml, a 15 ml polypropylene tube (Falcon 2059) is suitable for homogenization.

Tissues should be kept frozen until homogenized. The TRIzol is added directly to the frozen tissue before homogenization. Following homogenization, the insoluble material is removed from the homogenate by centrifugation at 7500 x g for 15 min. in a Sorvall superspeed or 12,000 x g for 10 min. in an Eppendorf centrifuge at 4°C. The cleared homogenate is then transferred to a new tube(s). Samples may be frozen and stored at -60 to -70°C for at least one month or else continue with the purification.

The next process is phase separation. The homogenized samples are incubated for 5 minutes at room temperature. Then, 0.2 ml of chloroform per 1ml of TRIzol reagent is added to the homogenization mixture. The tubes are securely capped and shaken vigorously by hand (do not vortex) for 15 seconds. The samples are then incubated at room temp. for 2-3 minutes and next centrifuged at 6500 rpm in a Sorvall superspeed for 30 min. at 4oC.

The next process is RNA Precipitation. The aqueous phase is transferred to a fresh tube. The organic phase can be saved if isolation of DNA or protein is desired. Then 0.5 ml of isopropyl alcohol is added per 1ml of TRIzol reagent used in the original homogenization. Then, the tubes are securely capped and inverted to mix. The samples are then incubated at room temp. for 10 minutes an centrifuged at 6500 rpm in Sorvall for 20 min. at 4°C.

The RNA is then washed. The supernatant is poured off and the pellet washed with cold 75% ethanol. 1 ml of 75% ethanol is used per 1 ml of the TRIzol reagent used in the initial homogenization. The tubes are capped securely and inverted several times to loosen pellet without vortexing. They are next centrifuged at <8000  $\,\mathrm{rpm}$  (<7500  $\,\mathrm{x}$  g) for 5 minutes at 4°C.

The RNA wash is decanted. The pellet is carefully transferred to an Eppendorf tube (sliding down the tube into the new tube by use of a pipet tip to help guide it in if necessary). Tube(s) sizes for precipitating the RNA depending on the working volumes. Larger tubes may take too long to dry. Dry pellet. The RNA is then resuspended in an appropriate volume (e.g., 2 -5 ug/ul) of DEPC H<sub>2</sub>0. The absorbance is then measured.

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The poly A+ mRNA may next be purified from total RNA by other methods such as Qiagen's RNeasy kit. The poly A + mRNA is purified from total RNA by adding the oligotex suspension which has been heated to 37°C and mixing prior to adding to RNA. The Elution Buffer is incubated at 70°C. If there is precipitate in the buffer, warm up the 2 x Binding Buffer at 65°C. The the total RNA is mixed with DEPC-treated water, 2 x Binding Buffer, and Oligotex according to Table 2 on page 16 of the Oligotex Handbook and next incubated for 3 minutes at 65°C and 10 minutes at room temperature.

The preparation is centrifuged for 2 minutes at 14,000 to 18,000 g, preferably, at a "soft setting," The supernatant is removed without disturbing Oligotex pellet. A little bit of solution can be left behind to reduce the loss of Oligotex. The supernatant is saved until satisfactory binding and elution of poly A+mRNA has been found.

Then, the preparation is gently resuspended in Wash Buffer OW2 and pipetted onto the spin column and centrifuged at full speed (soft setting if possible) for 1 minute.

Next, the spin column is transferred to a new collection tube and gently resuspended in Wash Buffer OW2 and centrifuged as described herein.

Then, the spin column is transferred to a new tube and eluted with 20 to 100 ul of preheated (70°C) Elution Buffer. The Oligotex resin is gently resuspended by pipetting up and down. The centrifugation is repeated as above and the elution repeated with fresh elution buffer or first eluate to keep the elution volume low.

The absorbance is next read to determine the yield, using diluted Elution Buffer as the blank.

Before proceeding with cDNA synthesis, the mRNA is precipitated before proceeding with cDNA synthesis, as components leftover or in the Elution Buffer from the Oligotex purification procedure will inhibit downstream enzymatic reactions of the mRNA. 0.4 vol. of 7.5 M NH4OAc + 2.5 vol. of cold 100% ethanol is added and the preparation precipitated at -20°C 1 hour to overnight (or 20-30 min. at -70°C), and centrifuged at 14,000-16,000 x g for 30 minutes at 4°C. Next, the pellet is wheshed with 0.5 ml of 80% ethanol (-20°C) and then centrifuged at 14,000-16,000 x g for 5 minutes at room temperature. The 80% ethanol wash is then repeated. The last bit of ethanol from the pellet is then dried without use of a speed vacuum and the pellet is then resuspended in DEPC H<sub>2</sub>0 at 1ug/ul concentration.

# Alternatively the RNA may be purified using other methods (e.g., Qiagen's RNeasy kit).

No more than 100 ug is added to the RNeasy column. The sample volume is adjusted to 100 ul with RNase-free water. 350 ul Buffer RLT and then 250 ul ethanol (100%) are added to the sample. The preparation is then mixed by pipetting and applied to an RNeasy mini spin column for centrifugation (15 sec at >10,000 rpm). If yield is low, reapply the flowthrough to the column and centrifuge again.

Then, transfer column to a new 2 ml collection tube and add 500 ul Buffer RPE and centrifuge for 15 sec at >10,000 rpm. The flowthrough is discarded. 500 ul Buffer RPE and is then added and the preparation is centriuged for 15 sec at >10,000 rpm. The flowthrough is discarded. and the column membrane dried by centrifuging for 2 min at maximum speed. The column is transferred to a new 1.5-ml collection tube. 30-50 ul of RNase-free water is applied directly onto column membrane. The column is then centrifuged for 1 min at >10,000 rpm and the elution step repeated.

The absorbance is then read to determine yield. If necessary, the material may be ethanol precipitated with ammonium acetate and 2.5X volume 100% ethanol.

#### 30 First Strand cDNA Synthesis

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The first strand can be make using Using Gibco's "SuperScript Choice System for cDNA Synthesis" kit. The starting material is 5 ug of total RNA or 1 ug of polyA+mRNAI. For total RNA, 2 ul of SuperScript RT is used; for polyA+mRNA, 1 ul of SuperScript RT is used. The final volume of first strand synthesis mix is 20 ul. The RNA should be in a volume no greater than 10 ul. The RNA is incubated with 1 ul of 100 pmol T7-T24 oligo for 10 min at 70°C followed by addition on ice of 7 ul of: 4ul 5X 1st Strand Buffer, 2 ul of 0.1M DTT, and 1 ul of 10mM dNTP mix. The preparation is then incubated at 37°C for 2 min before addition of the SuperScript RT followed by incubation at 37°C for 1 hour.

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#### Second Strand Synthesis

For the second strand synthesis, place 1st strand reactions on ice and add: 91 ul DEPC  $\rm H_20$ ; 30 ul 5X 2nd Strand Buffer; 3 ul 10mM dNTP mix; 1 ul 10 U/ul E.coli DNA Ligase; 4 ul 10 U/ul E.coli DNA Polymerase; and 1 ul 2 U/ul RNase H. Mix and incubate 2 hours at 16°C. Add 2 ul T4 DNA Polymerase. Incubate 5 min at 16°C. Add 10 ul of 0.5M EDTA.

## Cleaning up cDNA

The cDNA is purified using Phenol:Chloroform:Isoamyl Alcohol (25:24:1)
and Phase-Lock gel tubes. The PLG tubes are centrifuged for 30 sec at maximum speed.
The cDNA mix is then transferred to PLG tube. An equal volume of
phenol:chloroform:isamyl alcohol is then added, the preparation shaken vigorously (no
vortexing), and centrifuged for 5 minutes at maximum speed. The top aqueous solution is
transferred to a new tube and ethanol precipitated by adding 7.5X 5M NH4OAc and 2.5X
volume of 100% ethanol. Next, it is centrifuged immediately at room temperature for 20
min, maximum speed. The supernatant is removed, and the pellet washed with 2X with cold
80% ethanol. As much ethanol wash as possible should be removed before air drying the
pellet; and resuspending it in 3 ul RNase-free water.

### In vitro Transcription (IVT) and labeling with biotin

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In vitro Transcription (IVT) and labeling with biotin is performed as follows:

Pipet 1.5 ul of cDNA into a thin-wall PCR tube. Make NTP labeling mix by combining 2 ul

T7 10xATP (75 mM) (Ambion); 2 ul T7 10xGTP (75 mM) (Ambion); 1.5 ul T7 10xCTP (75

mM) (Ambion); 1.5 ul T7 10xUTP (75 mM) (Ambion); 3.75 ul 10 mM Bio-11-UTP

(Boehringer-Mannheim/Roche or Enzo); 3.75 ul 10 mM Bio-16-CTP (Enzo); 2 ul 10x T7

transcription buffer (Ambion); and 2 ul 10x T7 enzyme mix (Ambion). The final volume is

20 ul. Incubate 6 hours at 37°C in a PCR machine. The RNA can be furthered cleaned.

Clean-up follows the previous instructions for RNeasy columns or Qiagen's RNeasy protocol

handbook. The cRNA often needs to be ethanol precipitated by resuspension in a volume compatible with the fragmentation step.

Fragmentation is performed as follows. 15 ug of labeled RNA is usually fragmented. Try to minimize the fragmentation reaction volume; a 10 ul volume is recommended but 20 ul is all right. Do not go higher than 20 ul because the magnesium in 15 the fragmentation buffer contributes to precipitation in the hybridization buffer. Fragment RNA by incubation at 94 C for 35 minutes in 1 x Fragmentation buffer (5 x Fragmentation buffer is 200 mM Tris-acetate, pH 8.1; 500 mM KOAc; 150 mM MgOAc). The labeled RNA transcript can be analyzed before and after fragmentation. Samples can be heated to 65°C for 15 minutes and electrophoresed on 1% agarose/TBE gels to get an approximate idea of the transcript size ranse.

For hybridization, 200 ul (10 ug cRNA) of a hybridization mix is put on the chip. If multiple hybridizations are to be done (such as cycling through a 5 chip set), then it is recommended that an initial hybridization mix of 300 ul or more be made. The hybridization mix is: fragment labeled RNA (50 ng/ul final conc.); 50 pM 948-b control oligo; 1.5 pM BioB; 5 pM BioC; 25 pM BioD; 100 pM CRE; 0.1 mg/ml herring sperm DNA; 0.5 mg/ml acetylated BSA; and 300 ul with 1xMES hyb buffer.

The hybridization reaction is conducted with non-biotinylated IVT (purified by RNeasy columns) (see example 1 for steps from tissue to IVT): The following mixture is prepared:

IVT antisense RNA; 4 
$$\mu$$
g:  $\mu$ l Random Hexamers (1  $\mu$ g/ $\mu$ l): 4  $\mu$ l H<sub>2</sub>O:  $\mu$ l 14  $\mu$ l

5 Incubate the above 14 μl mixture at 70°C for 10 min.; then put on ice.

The Reverse transcription procedure uses the following mixture:

3 μΙ
0.6 µl
2.4 µl
3 μ1
1 μ1
16 11

Then above solution is added to the hybridization reaction and incubated for 30 min., 42°C.

Then, 1 ul SSII is added and incubated for another hour before being placed on ice.

The 50X dNTP mix contains 25mM of cold dATP, dCTP, and dGTP, 10mM of dTTP and is made by adding 25  $\mu$ l each of 100mM dATP, dCTP, and dGTP; 10  $\mu$ l of 100mM dTTP to 15  $\mu$ l H<sub>2</sub>O.]

RNA degradation is performed as follows. Add 86 µl H2O, 1.5 µl 1M NaOH/
2 mM EDTA and incubate at 65°C, 10 min.. For U-Con 30, 500 µl TE/sample spin at 7000 g
for 10 min, save flow through for purification. For Qiagen purification, suspend u-con
recovered material in 500 µl buffer PB and proceed using Qiagen protocol. For DNAse
digestion, add 1 ul of 1/100 dilution of DNAse/30 ul Rx and incubate at 37°C for 15 min.
Incubate at 5 min 95°C to denature the DNAse.

### Sample preparation

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For sample preparation, add Cot-1 DNA, 10  $\mu$ l; 50X dNTPs, 1  $\mu$ l; 20X SSC, 2.3  $\mu$ l; Na pyro phosphate, 7.5  $\mu$ l; 10 mg/ml Herring sperm DNA; 1  $\mu$ l of 1/10 dilution to 21.8 final vol. Dry in speed vac. Resuspend in 15  $\mu$ l  $\mu$ l  $\mu$ 0. Add 0.38  $\mu$ 1 10% SDS. Heat

95°C, 2 min and slow cool at room temp. for 20 min. Put on slide and hybridize overnight at 64°C. Washing after the hybridization: 3X SSC/0.03% SDS: 2 min., 37.5 ml 20X SSC+0.75ml 10% SDS in 250ml H<sub>2</sub>O; 1X SSC: 5 min., 12.5 ml 20X SSC in 250ml H<sub>2</sub>O; 0.2X SSC: 5 min., 2.5 ml 20X SSC in 250ml H<sub>2</sub>O. Dry slides and scan at appropriate PMT's and channels.

# Example 2: Taxol resistant Xenograft Model of Human Prostate Cancer

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Treatment regimens that include paclitaxel (Taxol; Bristol-Myers Squibb

Company, Princeton, NJ) have been particularly successful in treating hormone-refractory prostate cancer in the phase II setting (Smith et al., Semin. Oncol. 26(1 Suppl 2):109-11 (1999)). However, many patients develop tumors which are initially, or later become, resistant to taxol. To identify genes that may be involved with resistance to taxol, or are regulated in response to taxol resistance, and therefore may be used to treat, or identify, taxol resistance in patients, the following experiments were carried out.

The androgen-independent human cell line CWR22R was grown as a xenograft in nude mice (Nagabhushan et al., Cancer Res. 56(13):3042-3046 (1996); Agus et al., J. Natl. Cancer Inst. 91(21):1869-1876 (1999); Bubendorf et al., J. Natl. Cancer Inst. 91(20):1758-1764 (1999)). Initially, these xenograft tumors were sensitive to therapeutic doses of taxol. The mice were treated continuously with sub-therapeutic doses, and the tumors were allowed to grow for 3-4 weeks, before surgical removal of the tumors. The tumor from an individual mouse was then minced, and a small portion was then injected into a healthy nude mouse, establishing a second

passage of the tumor. This mouse was then treated continuously with the same sub-therapeutic dose of taxol. This process was repeated 14 times, and a portion of each generation of xenograft tumor was collected. There was increasing resistance to therapeutic doses of taxol with each generation. Bythe end of the process, the tumors were fully resistant to therapeutic doses of taxol. RNA from each generation of tumor was then isolated, and individual mRNA species were quantified using a custom Affymetrix GeneChip® oligonucleotide microarray, with probes to interrogate approximately 35,000

unique mRNA transcripts. Genes were selected that showed a statistically significant upregulation, or down-regulation, during the subsequent generations of increasingly taxolresistant tumors. Only one gene was significantly up-regulated, whereas 24 genes were down-regulated; these are presented in Table 10.

The gene sequences identified to be overexpressed in prostate cancer may be used to identify coding regions from the public DNA database. The sequences may be used to either identify genes that encode known proteins, or they may be used to predict the coding regions from genomic DNA using exon prediction algorithms, such as FGENESH (Salamov and Solovyev, 2000, Genome Res. 10:516-522). In addition, one of ordinary skill in the art would understand how to obtain the unigene cluster identification and sequence information according to the exemplar accession numbers provided in Tables 1-16. (see,

10 http://www.ncbi.nlm.nih.gov/UniGene/).

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TABLE1: shows genes, including expression sequence tags, differentially expressed in prostate tumor tissue compared to normal tissue as analyzed using the Affymetrix/Bos Hu01 GeneChip array. Shown are the relative amounts of each gene expressed in prostate tumor samples and various normal tissue samples showing the highest expression of the gene.

Pkey: Unique Eos probeset identifier number
Exemplar Accession number, Genbank accession number
Unigenen Title:
Unigene Title:
Rit: Ratio of tumor to normal body tissue

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	Pkey	UnigenelD	ExAccn	Uningene Title	R1
	131010	Hs.272458	AA121266	ESTs	37.2
		Hs.290905		ESTs: Weakly similar to (defline not ava	32.6
20		Hs.31412		ESTS	30.1
20		Hs.1852	M24902	acid phosphatase; prostate	25.2
		Hs.279477		FSTs	24.8
		Hs.183752		microseminoprotein; beta-	23.8
		Hs.171995		kallikrein 3; (prostate specific antigen	21.4
25		Hs.57771		Homo sapiens mRNA for serine protease (T	18.9
23		Hs.162859		ESTs	18.6
		Hs.30343		ESTs	17.4
		Hs.1832	K01911	neuropeptide Y	17.3
		Hs.1915	N48056	folate hydrolase (prostate-specific memb	17
30		Hs.40808		ESTs	16.9
		Hs.262476		S-adenosylmethionine decarboxylase 1	16.7
		Hs.33287		ESTs	16.5
		Hs.11260		ESTs	16.4
				Antigen, Prostate Specific, Alt. Splice	16
35		Hs.181350		kallikrein 2; prostatic	15.4
		Hs.99872		fetal Alzheimer antigen	15
		Hs.62192	M27436	coagulation factor ill (thromboplastin;	13.9
		Hs.8236	D62633	ESTs	12.7
	133944	Hs.7780	AA045870	ESTs	12.5
40	109141	Hs.193380	AA176428	ESTs	12.3
	130974	Hs.2178	X57985	H2B histone family; member Q	11.8
	114768	Hs.182339	AA149007	ESTs	11.8
	104394	Hs.172129	H46617	yp19h1.r1 Soares breast 3NbHBst Homo sap	11.8
		Hs.102720		ESTs	11.6
45	104660	Hs.14846	AA007160	ESTs	11.4
	100116	Hs.78045	D00654	actin; gamma 2; smooth muscle; enteric	11
	131061	Hs.268744	N64328	ESTs; Moderately similar to KIAA0273 [H.	10.9
	126645	126645	Al167942	Homo saplens BAC done RG041D11 from 7q2	10.7
	135153	Hs.95420	N40141	Homo sapiens mRNA for JM27 protein; comp	10.6
50	107033	Hs.113314	AA599629	ESTs	10.6
	118417		N66048	ESTs; Weakly similar to polymerase [H.sa	10.5
	126758	Hs.293960	W37145	ESTs	10.2
	115874	Hs.8364	AA406542	ESTs	10.1
	134989	Hs.92381	AA236324	ESTs; Weakly similar to IIII ALU CLASS A	10.1
55	107102	Hs.30652	AA609723	ESTs	10.1
		Hs.15641		ESTs	10.1
		Hs.59822		ESTs	10
		Hs.203270		ESTs	9.9
		Hs.121017		H2A histone family; member A	9.8
60		Hs.83663		ESTs	9.7
		Hs.80296		Purkinje cell protein 4	9.7
	117984	Hs.106778	N51919	ESTs	9.7
		Hs.22209		ESTs	9.4
		Hs.274509		T-cell receptor; garrima cluster	9.4
65		Hs.167133		ESTs	9.2
	121853	Hs.98502	AA425887	ESTs	9

8.8

		Hs.91011 Hs.55999	AA421562 W47380	anterior gradient 2 (Xenopus laevis; sec	8.9 8.9
				Protein Kinase Ht31, Camp-Dependent	8.9
		Hs.23317	AA281245	ESTs	8.8
5		Hs.76422	M22430	phospholipase A2; group IIA (platelets;	8.7
		Hs.31146	AA456264	ESTs; Highly similar to (defline not ava	8.5
		Hs.293185		yz61c5.s1 Soares_multiple_sclerosis_2NbH	8.5
		Hs.49397	N67869	ESTs	8.2 8.2
10		Hs.76704 Hs.334762	T68510	ESTs ESTs; Wealdy similar to KIAA0319 [H.sapi	8.1
10		Hs.20415	AAU000002 AA402000	ESTs; Weakly similar to GS3786 [H.sapien	8
		Hs.278695		ESTs	8
		Hs.66052	D84276	CD38 antigen (p45)	8
	114132	Hs.24192	Z38688	ESTs	7.9
15		Hs.301527		tumor necrosis factor (figand) superfami	7.7
		Hs.23023	AA456135	ESTs	7.8
		Hs.105700		secreted frizzled-related protein 4	7.5 7.4
		Hs.72472 Hs.22627	AA250737 R43162	ESTs ESTs	7.1
20	102398	NS.22021	U42359	Human N33 protein form 1 (N33) gene, exo	7
20	101201	Hs.2256	L22524	matrix metalloproteinase 7 (matrilysin;	6.9
	109272	Hs.288462		ESTs	6.9
		Hs.169849		myosin-binding protein C; slow-type	6.9
		Hs.155891		pre-B-cell leukemia transcription factor	6.8
25		Hs.302267		ESTs; Weakly similar to W01A6.c [C.elega	6.8 6.8
		Hs.257924 Hs.326416		ESTs ESTs	6.7
		Hs.173684		ESTs; Weakly similar to (defline not ava	6.7
		Hs.171995		kalikrein 3; (prostate specific antigen	6.6
30		Hs.26691	AA219134	ESTs	6.6
		Hs.18193	AA281591	Homo sapiens mRNA; cDNA DKFZp586B211 (fr	6.6
		Hs.59838	AA490969	ESTs	6.6
		Hs.323378		H.sapiens mRNA for transmembrane protein	6.6
35		Hs.75746	U07919	aldehyde dehydrogenase 6 ESTs; Moderately similar to APXL gene pr	6.5 6.5
33	100343	Hs.278628 Hs.108787	741050	Homo sepiens Mod4p homolog mRNA; complet	6.5
		Hs.126085		ESTs	6.5
		Hs.3383	AA010163	upstream regulatory element binding prot	6.5
	133376	Hs.7232	T23670	ESTs	8.4
40		Hs.8768	AA236559	ESTs; Weakly similar to neuronal thread	6.4
	104674	Hs.26289	AA009527	ESTs	6.4
	100/27	Hs.334786 Hs.15113	AF000573	Human HF.12 gene mRNA homogentisate 1;2-dioxygenase (homogenti	6.3
		Hs.278428		Homo sapiens mRNA for KIAA0896 protein;	6.3
45		Hs.250528		ESTs; Weakly similar to ANKYRIN; BRAIN V	6.3
		Hs.296638		prostate differentiation factor	6.3
	116429	Hs.279923		ESTs; Weakly similar to similar to GTP-b	6.2
		Hs.878	L29008	sorbitol dehydrogenase	6.2
£0.		Hs.37744	AA011176	ESTS	8.2 6.2
50	127248	Hs.179902	AA325029	EST27953 Cerebellum II Homo sapiens cDNA ESTs: Weakly similar to (defline not ava	6.2
	105500	Hs.222399	AA256485	ESTs	6.1
		Hs.2714	X74142	forkhead (Drosophila)-like 1	- 6.1
		Hs.40289	AA234767	ESTs	6
55		Hs.203213		ESTs	5.9
			AA281793	ESTs	5.8
		Hs.301997		ESTs	5.7 5.7
		Hs.48948	AA491457	ESTs ESTs	5.7
60		Hs.61539 Hs.125019	AA034020 738830	ESTs; Weakly similar to IIII ALU SUBFAMI	5.6
00		Hs.289072		ESTs	5.6
		Hs.170195		bone morphogenetic protein 7 (osteogenic	5.6
	124777	Hs.140237	R41933	ESTs; Weakly similar to neuronal thread	5.6
		Hs.337616		phosphodiesterase 3B; cGMP-inhibited	5.6
65		Hs.62354	M83822	Human beige-like protein (BGL) mRNA; par	5.5 5.5
		Hs.45107	N41002	ESTs heat shock 70kD protein 1	5.5 5.5
	132387	Hs.281434 Hs.98732	AA431407	Homo sapiens Chromosome 16 BAC clone CIT	5.5
		Hs.282476		S-adenosylmethionine decarboxylase 1	5.5

	113938		W81598	ESTs	5.4
		Hs.246315		ESTs	5.4
		Hs.75722	Al283493	ribophorin II	5.4
~			T34527	UDP-N-acetyl-a/pha-D-galactosamine:potyp	5.4 5.3
5		Hs.7780	AA056482	ESTs	5.3
		Hs.21223 Hs.328392	D17408	calponin 1; basic; smooth muscle	5.3
		Hs.98944	AA365031	Human guanine nucleotide exchange factor ESTs	5.3
		Hs.167531		ESTs; Weakly similar to (defline not ava	5.3
10		Hs.106336		ESTS: Weakly similar to IIII ALU SUBFAMI	5.3
10		Hs.25351	U90304	iroguois-class homeodomain protein	5.3
		Hs.194369		Homo sapieris chromosome 1 atrophin-1 rel	5.3
		Hs.109201		ESTs; Highly similar to (defline not ava	5.2
		Hs.79428	U15174	BCL2/adenovirus E18 19kD-interacting pro	5.2
15		Ha.159872		ESTs	5.2
	104787		AA027317	ESTs; Weakly similar to !!!! ALU SUBFAMI	5.2
	123527	Hs.108327	AA608679	damage-specific DNA binding protein 1 (1	5.2
	116646	Hs.194228	F03048	ESTs; Moderately similar to fill ALU SUB	5.2
		Hs.195850		keratin 5 (epidermolysis bullosa simplex	5.1
20		Hs.184598		ESTs; Weakly similar to IIII ALU SUBFAMI	5.1
		Hs.281428		ESTs; Moderately similar to !!!! ALU SUB	5.1
		Hs.169119		ESTs	5.1
		Hs.54416	X91868	sine oculis homeobox (Drosophila) homolo	5.1
05		Hs.106778		ESTs; Highly similar to (defilne not ava	5.1
25		Hs.148932		ESTs; Moderately similar to semaphorin V ESTs	5.1 5.1
		Hs.226434 Hs.47144	AA479362	ESTS	5
		Hs.80342	X07696	keratin 15	5
		Hs.326035		early growth response 1	5
30	126023	113.00.000	H58881	vr36d09.r1 Scares fetal liver spieen 1NF	5
50		Hs.13804	AA037316	ESTs	5
		Hs.39288	N93839	ESTs; Weakly similar to IIII ALU SUBFAMI	5
		Hs.89732	X78932	zinc finger protein 273	5
		Hs.23311	AB002365	Human mPNA for KIAA0367 gene; partial od	4.9
35	132047	Hs.3796	D83492	EphB6	4.9
	132880	Hs.177537	AA444369	ESTs	4.9
		Hs.74519	F10523	primase; polypeptide 2A (58kD)	4.8
		Hs.71119	U42360	Human N33 mPNA; complete cds	4.8
	104776		AA026349	ESTs	4.8
40		Hs.128749		Homo saplens alpha-methylacyl-CoA racema	4.8 4.8
		Hs.143067		ESTs Homo sapiens mRNA for KIAA0860 protein;	4.8
		Hs.26009 Hs.3585	AA233168	ESTs: Weakly similar to coded for by C.	4.8
		Hs.284186		ESTs vieakly similar to coded for by C.	4.8
45		Hs.183390		ESTs; Weakly similar to ZINC FINGER PROT	4.8
7.7		Hs.288126		ESTS	4.8
	125982	H3.200120	R98091	RAE1 (RNA export 1; S.pombe) homolog	4.8
		Hs.26243	W44682	ESTs	4.8
		Hs.117950		multifunctional polypeptide similar to S	4.7
50	129735		W80701	ESTs; Weakly similar to HERV-E envelope	4.7
	104479	Hs.106390	N36040	ESTs	4.7
	103731		AA070545	zm7c3.r1 Stratagene neuroepithelium (#93	4.7
	126575	Hs.127602	W72416	ESTs -	4.7
		Hs.231500		Human glucose transporter like protein-l	4.7
55	130617	Hs.1674	M90516	glutamine-fructose-6-phosphate transamin	4.7
		Hs.91622	H06373	Homo sapiens clone 24456 mRNA sequence	4.7 4.7
		Hs.82007	D42084	Human mRNA for KiAA0094 gene; partial cd	4.7
		Hs.32990	Al479264 AA610086	ESTs ESTs	4.7
60	131838	Hs.239489		TIA1 cytoloxic granule-associated RNA-bi	4.7
00	1145/2	Hs.91011	AA055788	ESTs	4.6
	103806	110.01011	AA130614	zo1f2.r1 Stratagene neuroepithelium NT2R	4.8
	130529		AA173238	small inducible cytokine A5 (RANTES)	4.6
		Hs.82065	AA406546	ESTs	4.6
65	111386	Hs.293798		ESTs	4.6
-		Hs.29679	AA452411	ESTs	4.6
	119943	Hs.14158	W86835	copine III	4.6
		Hs.100070		EST	4.6
	100774	Hs.89603	HG371-HT1063	Much 1, Epithelial, Alt. Splice 6	4.6

				Ret Transforming Gene	4.6
	132015	Hs.3731	D11900	ESTS	4.6 4.6
	126086	Hs.173094	H70975	yr/3g01.r1 Soares fetal liver spieen 1NF ESTs	4.6
5	108390	Hs.20166	AA446964	Prostate stem cell antigen	4.6
-	126959	110.00100	AA199853	ESTs: Moderately similar to IIII ALU SUB	4.5
		Hs.29117	X91648	H.saplens mRNA for pur alpha extended 3	4.5
		Hs.20953	AA039481	ESTs	4.5
10	125661		R50319	ESTs	4.5
10		Hs.234726 Hs.199160		alpha-1-antichymotrypsin ESTs	4.5 4.5
		Hs.75730		signal recognition particle receptor ('d	4.5
				Small Nuclear Ribonucleoprotein U1, 1snr	4.5
		Hs.7956	AA425906	ESTs	4.5
15		Hs.317584		ESTs	4.5
		Hs.24758		ESTs	4.5
		Hs.44566	U28831 T89386	Human protein immuno-reactive with anti-	4.4
	132056	Hs.38176 Hs.198760		Homo sapiens mRNA for KIAA0606 protein; neurofilament; heavy polypeptide (200kD)	4.4
20		Hs.1848	M22898	tumor protein p53 (Li-Fraumeni syndrome)	4.4
20		Hs.284296		ESTs; Highly similar to surface 4 integr	4.4
		Hs.22514	AA383142	ESTs	4.4
		Hs.119394		ESTs	4.4
25		Hs.29894	N79565	ESTs	4.4
25		Hs.98518	AA429278	ESTs	4.4 4.4
		Hs.211577 Hs.288989		ESTs; Highly similar to CG1 protein [H.s ESTs; Weakly similar to !!!! ALU SUBFAMI	4.4
		Hs.323966		ESTs: Moderately similar to IIII ALU SUB	4.4
		Hs.21941	AA187490	ESTs	4.3
30	127315		AA640834	nr27b06.r1 NCI_CGAP_Pr3 Homo saplens cDN	4.3
		Hs.54424	X87870	H.saplens mRNA for hepatocyte nuclear fa	4.3
		Hs.282990		ESTs; Weakly similar to F52C12.2 [C.eleg FST	4.3 4.3
		Hs.47567 Hs.278427		carebellar degeneration-related protein	4.3
35		Hs.114688		ESTs	4.3
33		Hs.105130		EST	4.3
	109391	Hs.184245		ESTs	4.3
	109175		AA180496	ESTs	4.3
40	127003	Hs.173540	AA550806	ESTs; Weakly similar to (defline not ava	4.3 4.3
40		Hs.46638 Hs.79993	U88871	chromosome 11 open reading frame 8 peroxisomal biogenesis factor 7	4.3
		Hs.5462	AF007216	solute carrier family 4; sodium bicarbon	4.3
			AA094720	ESTs; Weakly similar to (defline not ava	4.3
		Hs.295923		seven in absentia (Orosophila) homolog 1	4.3
45		Hs.93872		ESTs	4.3
		Hs.334762		ESTs; Weakly similar to KIAA0319 [H.sapi	4.2 4.2
		Hs.98747 Hs.6574	AA431732 AF007165	EST suppressin (nuclear deformed epidermal a	4.2
		Hs.20843	H04649	ESTs	4.2
50		Hs.69997	R79723	H,sapiens mRNA for transiin associated z	4.2
	134436	Hs.83190	S80437	fatty acid synthase (3' region) [human,	4.2
		Hs.251064		NBR2	4.2
		Hs.27413	AA436158	ESTs	- 4.2 4.2
55		Hs.248210 Hs.59815	X55777 W99362	H.sapiens Mahlavu hepatocellular cardino EST	4.2
33		Hs.283978		ESTs; Highly similar to (defline not ava	4.2
		Hs.1179	D90359	TATA box binding protein (TBP)-associate	4.2
	106586		AA455921	ESTs; Weakly similar to IIII ALU SUBFAMI	4.2
		Hs.29852	R79220	ESTs	4.2
60		Hs.279929		H.sapiens mRNA for gp25L2 protein	4.2 4.2
		Hs.57419 Hs.326292	U25435	transcriptional repressor ESTs	4.2
		Hs.94109	AA489046	ESTs	4.2
		Hs.105938		lactotransferrin	4.1
65	129133	Hs.108850	R56728	yg95c6.r1 Soares Infant brain 1NIB Homo	4.1
		Hs.6641	N98707	kinesin family member 5C	4.1
		Hs.14051	AA351779	ESTs	4.1 4.1
		Hs.45032 Hs.327179	AA192157	ESTs solute carrier family 17 (sodium phospha	4.1
	10/3/0	110,02/1/9	00000	solute carrier taring 17 (account prospins	4.1

	128517	Hs.100861	AA280617	ESTs; Weakly similar to p60 katanin [H-s	4.1
	130555	Hs.116774		ESTs	4.1
		Hs.24183	AA343514	ESTs	4.1
5		Hs.26369 Hs.181889	AA133237	ESTs ESTs	4.1 4.1
J		Hs.172129		ESTs; Moderately similar to lit! ALU SUB	4.1
		Hs.3085	D29677	KIAA0054 gene product	4.1
		Hs.118127		actin; alpha; cardiac muscle	4.1
		Hs.12913		ESTs; Weakly similar to (defline not ava	4.1
10		Hs.8859	AA128486	ESTs	4.1
	126735	Hs.226795	AA808949	glutathione S-transferase pi	4.1
		Hs.8036	T26471	ESTs; Moderately similar to IIII ALU SUB	4
	102460	Hs.211582	U48959	Homo sapiens myosin light chain kinase (	4
16		Hs.26813		ESTs; Weakly similar to (defline not ava	4
15		Hs.104207 Hs.287987		ESTs: Weakly similar to IIII ALU SUBFAMI	4
			AA234561	ESTS, Weakly similar to the ALO SOSPAMI	
		Hs.42736	AA291946	ESTs	4
		Hs.97293	AA293656	ESTS	4
20		Hs.94560	726317	desmogleln 2	4
		Hs.144941	Al147408	ESTs	4
		Hs.25320	AA447223	ESTs	4
	128046		AA873285	ESTs	4
		Hs.114366		pyrroline-5-carboxylate synthetase (glut	4
25			AA449455	ESTs	4
		Hs.86278	W27601 AA487015	ESTs; Moderately similar to (define not	4 3.9
	129593	Hs.98314 Hs.31608	H18836	ESTs; Weakly similar to !!!! ALU SUBFAMI ESTs	3.9
		Hs.8645	AA235303	ESTS	3.9
30	104791			ESTs	3.9
50		Hs.111496		ESTs	3.9
	127800	Hs.79428	AA521047	BCL2/adenovirus E1B 19kD-Interacting pro	3.9
		Hs.167904		ESTs	3.9
~ "		Hs.163960		ESTs	3.9
35		Hs.198726		vasoactive Intestinal peptide receptor 1	3.9
		Hs.75216	Y00615	protein tyrosine phosphatase; receptor t caldesmon 1	3.9 3.9
		Hs.325474 Hs.301985		ESTs	3.9
		Hs.81086	AA460012	solute carrier family 22 (organic cation	3.9
40		Hs.50421	R38102	KIAA0203 gene product	3.9
••		Hs.241493		natural killer-tumor recognition sequenc	3.9
	103695	Hs.186600	AA018758	ESTs	3.9
				Caldesmon 1, Alt. Splice 6, Non-Muscle	3.9
		Hs.78771	D82614	ESTs	3.9
45		Hs.19978	H26417	ESTs	3.9 3.9
	125298			ESTs zt87a9.r1 Soares_testis_NHT Homo sapiens	3.9
	104060 105823			ESTs	3.9
	126499			ESTs; Moderately similar to unknown [M.m	3.9
50		Hs.18895	D50927	KIAA0137 gene product	3.8
	123494		AA599786	ESTs	3.8
	104846	Hs.32478	AA040154	ESTs	3.8
		Hs.71721	AA142913	ESTs	- 3.8
		Hs.45207	AA292537	ESTs	3.8
55		Hs.241552		Human mRNA for KIAA0268 gene; partial cd	3.8 3.8
		Hs.129228 Hs.102859		galactokinase 2 ESTs	3.8
		Hs.24427	AA247788	ESTs; Highly similar to (delline not ava	3.8
		Hs.269228		ESTs	3.8
60		Hs.73848		ESTs	3.8
		Hs.9394	AA495926	ESTs	3.8
		Hs.620	M69225	bullous pemphigoid antigen 1 (230/240kD)	3.8
		Hs.14912	AA424524	Homo sapiens mRNA for KiAA0286 gene; par	3.8
		Hs.269721		ESTs	3.8
65		Hs.58694	W92051	ESTs zh98g04.r1 Scares_fetal_liver_spleen_1NF	3.8 3.8
		Hs.50382 Hs.112969	AA007489 AA621311	EST	3.7
		Hs.17752	H95978	Homo sapiens phosphatidy/serine-specific	3.7
		Hs.162	M35410	Insulin-like growth factor binding prote	3.7

	447007	No. 44700	NOCOLA	ser-Thr protein kinase related to the my	3.7
		Hs.44708	N39214	Sel-Trif protein Milase relation to the my	
		Hs.39712		ESTs; Weakly similar to BONE/CARTILAGE P	3.7
	100379	Hs.278721	D82060	Homo saplens mRNA for membrane protein w	3.7
	115646	Hs.305971	AA404352	ESTs	3.7
5		Hs.193700		ESTs; Moderately similar to IIII ALU SUB	3.7
,					3.7
	102162	Hs.1592	U18291	CDC16 (cell division cycle 16; S. cerevi	
	128530	Hs.183475	AA504343	ESTs; Moderately similar to IIII ALU SUB	3.7
	119940	Hs.272531	W86779	EST	3.7
		Hs.23837		yw34b06.s1 Morton Fetal Cochlea Homo sap	3.7
10	110/08	115.23007	A L LOCOCOT		3.7
10	132914	Hs.60293	AA496037	ESTs	
	113594	Hs.15663	T92030	ESTs	3.7
	103702	Hs,279952	AA027793	ESTs; Highly similar to (defline not ava	3.7
	100700	Hs.19347	A AD 40 400	ESTs	3.7
	130700	ns.1934/	MM2404U0		3.7
		Hs291025		EST	
15	120691	Hs.22380	AA291173	ESTs	3.7
	103153	Hs.75295	X68534	guanylate cyclase 1; soluble; alpha 3	3.7
		Hs.109390		ESTs	3.7
				ESTs	3.7
		Hs.54900			
		Hs.7337	AA512902	ESTs	3.7
20	105503	Hs.31707	AA256616	ESTs	3.7
	104280	Hs.194283	AF008192	Homo sapiens putative GR6 protein (GR6)	3.7
		Hs.35699		ESTs	3.7
					3.6
		Hs.105273		ESTs	
	103862	Hs.6363	AA206625	ESTs	3.6
25	100696	Hs.121686	HG3162-HT3339	Transcription Factor lia	3.6
		Hs.166994		FAT tumor suppressor (Drosophila) homolo	3.6
		110.10000	Y10511	H.sapiens mRNA for CD176 protein	3.6
	103520				
		Hs.302738		ESTs	3.6
	101638	Hs.75511	M92934	connective tissue growth factor	3.8
30	113702		T97307	ESTs; Moderately similar to !!!! ALU SUB	3.6
		Hs.48426		EST	3.6
		Hs.68554		EST	3.6
					3.6
			AA400517	ESTs; Moderately similar to UDP-GLUCOSE:	
		Hs.170291		ESTs	3.6
- 33	127858	Hs.27973	AA806365	oc26h07.s1 NCL_CGAP_GCB1 Homo saplens cD	3.6
35		Hs.27973		oc26h07.s1 NCL_CGAP_GCB1 Homo saplens cD dioxin-responsive pene (putative polyade	3.6
33	101964		S81578	dioxin-responsive gene (putative polyade	3.6
33	101964 105508	Hs.326416	S81578 AA256680	dioxin-responsive gene (putative polyade ESTs	3.6
33	101964 105508 116844	Hs.326416 Hs.337434	S81578 AA256680 H64938	dioxin-responsive gene (putative polyade ESTs ESTs	3.6 3.6 3.6
	101964 105508 116844 105372	Hs.326416 Hs.337434 Hs.142296	S81578 AA256680 H64938 AA236481	dioxin-responsive gene (putative polyade ESTs ESTs ESTs	3.6 3.6 3.6
40	101964 105508 116844 105372	Hs.326416 Hs.337434 Hs.142296	S81578 AA256680 H64938 AA236481	dioxin-responsive gene (putative polyade ESTs ESTs ESTs	3.6 3.6 3.6 3.8
	101964 105508 116844 105372 100745	Hs.326416 Hs.337434 Hs.142296 Hs.144630	\$81578 AA256680 H64938 AA236481 HG3510-HT3704	dioxin-responsive gene (putative polyade ESTs ESTs ESTs V-Erba Related Ear-3 Protein	3.6 3.6 3.6 3.8
	101964 105508 116844 105372 100745 127521	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018	\$81578 AA256680 H64938 AA236481 HG3510-HT3704 AA809962	diodin-responsive gene (putative polyade ESTs ESTs ESTs V-Erba Related Ear-3 Protein ESTs	3.6 3.6 3.6 3.6 3.8
	101964 105508 116844 105372 100745 127521 110758	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018 Hs.274265	\$81578 AA256680 H64938 AA236481 HG3510-HT3704 AA809962 N21365	dioxim-responsive gene (putative polyade ESTs ESTs ESTs V-Ends Related Ear-3 Protein ESTs	3.6 3.6 3.6 3.8 3.6 3.6
	101964 105508 116844 105372 100745 127521 110758 107307	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018 Hs.274265 Hs.44155	\$81578 AA256680 H64938 AA236481 HG3510-HT3704 AA809962 N21365 T52069	dioxin-responsive gene (putative polyade ESTs ESTs ESTs ESTs V-Etha Related Ear-3 Protein ESTs talin constinct kinase; milochondrial 2 (sarcom	3.6 3.6 3.6 3.8 3.6 3.6 3.6
40	101964 105508 116844 105372 100745 127521 110758 107307 133200	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018 Hs.274265 Hs.44155 Hs.183639	\$81578 AA256680 H64938 AA236481 HG3510-HT3704 AA809962 N21385 T52069 AA432248	diodri-responsive gene (putative polyacie EST e EST e EST e V-Erba Related Ear-3 Protein EST s Latin creatine kinase; milochondrial 2 (sarcom EST s	3.6 3.6 3.6 3.8 3.6 3.6 3.6 3.6
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40	101964 105508 116844 105372 100745 127521 110758 107307 133200 114774	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018 Hs.274265 Hs.44155 Hs.183639 Hs.184325	\$81578 AA256680 H64938 AA238481 HG3510-HT3704 AA809962 N21335 T52099 AA432248 AA150343	diodn-responsive gene (putative polyade ESTE ESTE ESTE VE-fina Related Ear-3 Protein EST talin constant (knase; mitochondrial 2 (sarcom ESTE	3.6 3.6 3.6 3.6 3.6 3.6 3.6 3.6
40	101964 105508 116844 105372 100745 127521 110758 107307 133200 114774 120265	Hs.326416 Hs.337434 Hs.142296 Hs.144630 Hs.164018 Hs.274265 Hs.44155 Hs.183639 Hs.184325 Hs.270696	\$81578 AA255680 H64938 AA236481 HG3510-HT3704 AA809962 N21385 T52069 AA432248 AA150043 AA173759	diodn-responsive gene (putative polyacie ESTe ESTe ESTin ESTin ESTin ESTin ESTin ESTin ESTin ESTin ESTin ESTin Hoderately similar to IIII ALU SUB	3.6 3.6 3.6 3.8 3.6 3.6 3.6 3.6 3.6
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40 45 50 55 60	101984 105508 10	Hs.328416 Hs.337434 Hs.1442296 Hs.1446301 Hs.164018 Hs.124235 Hs.274235 Hs.193067 Hs.48529 Hs.270996 Hs.270999 Hs.35941 Hs.279780 Hs.35941 Hs.78793 Hs.334641 Hs.79572 Hs.361804 Hs.17921 Hs.28178 Hs.303815 Hs.262797 Hs.26279 Hs.303815 Hs.303815 Hs.26279 Hs.303815 Hs.	\$41579 \$412594	doubn-responsive jens (putalive polyade EST2 EST2 EST3 EST3 EST3 EST3 EST3 EST3 EST3 EST3	3.6. 3.6. 3.6. 3.6. 3.6. 3.6. 3.6. 3.6.
40 45 50 55 60	101984 (10570 116844 (10570 116844 (10570 116844 (10570 116844 (10570 116844 (10570 11684 (10780 11684 (10780 11684 (10780 11684 (10880	Ha 326416 Ha 327434 Ha 14226 Ha 144630 Ha 164618 Ha 144630 Ha 16418 Ha 18363 Ha 184325 Ha 184325	\$41579 AA259690 H64938 H64938 H64938 H64938 H74939 H74939 AA25949 AA15996 AA45924 AA15949 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430	doubn-responsive jens (putalive polyade ESTE ESTE ESTE ESTE ESTE ESTE ESTE EST	3.6.6 3.6 3
40 45 50 55 60	101984 (10570 116844 (10570 116844 (10570 116844 (10570 116844 (10570 116844 (10570 11684 (10780 11684 (10780 11684 (10780 11684 (10880	Hs.328416 Hs.337434 Hs.1442296 Hs.1446301 Hs.164018 Hs.124235 Hs.274235 Hs.193067 Hs.48529 Hs.270996 Hs.270999 Hs.35941 Hs.279780 Hs.35941 Hs.78793 Hs.334641 Hs.79572 Hs.361804 Hs.17921 Hs.28178 Hs.303815 Hs.262797 Hs.26279 Hs.303815 Hs.303815 Hs.26279 Hs.303815 Hs.	\$41579 AA259690 H64938 H64938 H64938 H64938 H74939 H74939 AA25949 AA15996 AA45924 AA15949 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430 M5430	doubn-responsive jens (putalive polyade EST2 EST2 EST3 EST3 EST3 EST3 EST3 EST3 EST3 EST3	3.6. 3.6. 3.6. 3.6. 3.6. 3.6. 3.6. 3.6.

	102663	Hs.168075	U70322	karyopherin (importin) beta 2	3.5
	120240	Hs.13531	AA442868	ESTs: Weakly similer to (define not ava.	3.5
	132154	Hs.41119	N67179	ESTs	3.5
	101000	Hs.30695	AA599653	transcription factor-like 5 (basic helix	3.5
~					
5	127862	Hs.163191	AA765305	EST	3.5
	100000	Hs.189810	Whenen	Human DNA sequence from PAC 388M5 on chr	3.5
		118, 1090 10			3.0
	119071		R31180	ESTs	3.5
		Hs.96593	AA282978	ESTs	3.5
	110721	Hs.31319	H97678	ESTs	3.5
10		Hs.43086	AA011247	ESTs	3.5
10					
	103106	Hs.1857	X62025	phosphodiesterase 6G; cGMP-specific; rod	3.5
		Hs.90797	AA504806	Harris and lane alone 00000 mONA assurance	3.5
				Homo sapiens clone 23620 mRNA sequence	
	105309	Hs.4104	AA233790	ESTs	3.5
					3.5
		Hs.19525	R39390	ESTs	
15	100101	Hs.52184	AA167708	ESTs	3.5
13					
	103134	Hs.2839	X65724	Nortie disease (pseudoglioma)	3.5
	191700	Hs.301449	YRROOR	adenovirus 5 E1A binding protein	3.5
	118535	Hs.49418	N67968	ESTs	3.5
	100500	Hs.11223	U62389	Human putative cytosolic NADP-dependent	3.4
20	125905	Hs.6456	T69868	chaperonin containing TCP1; subunit 2 (b	3.4
	100100	Hs.301997	AA470207	ESTs	3.4
	105327	Hs.211593	AA234440	ESTs	3.4
		Hs.57787	AA456598	ESTs	3.4
		U8'01101			
	122635		AA454085	EST	3.4
25		Hs.260116			3.4
43				metalloprotease 1 (pitrilysIn family)	
	121038	Hs.34956	AA283620	ESTs	3.4
				ESTs	3.4
	1338/1	Hs.182793	WW404081		
	107175	Hs.292503	44821751	ESTs: Weakly similar to KIAA0601 protein	3.4
					3.4
	101188	Hs.184298	120320	cyclin-dependent kinase 7 (homolog of Xe	
30	126422	Hs.237658	H48518	ESTs; Highly similar to apolipoprotein A	3.4
50		110.201000		COT-, March and a to the ALL OLACO D	3.4
	118475		N66845	ESTs; Weakly similar to !!!! ALU CLASS B	
	104558	Hs.88959	R56678	ESTs; Weakly similar to !!!! ALU SUBFAMI	3.4
	128307	Hs.132005	Al453794	ESTs	3.4
	110054	Hs.25829	R51831	ESTs	3.4
0.0					
35	125408	Hs,89578	N72353	yv37e12.r1 Soares fetal liver spieen 1NF	3.4
	100024	Hs.175955	MOOROA	ESTs	3.4
	130844	Hs.20191	D12122	seven in ebsentia (Drosophila) homolog 2	3.4
	107149	Hs.20843	AA533553	ni68h04.s1 NCL CGAP_Pr10 Homo sapiens cD	3.4
	135309	Hs.42500	D25984	ESTs	3.4
40	100704	Hs.295978	A A 002407	stimulated trans-acting factor (50 kDa)	3.4
40					
	127892	Hs.187983	Al021912	ESTs	3.4
		Hs.92127	F04816	ESTs	3.4
	134700	Hs.8868	AA481414	golgi SNAP receptor complex member 1	3.4
		Hs.166196	A 400 4000	ESTs	3.4
45	103649	Hs.155963	Z70219	H.sapiens mRNA for 5'UTR for unknown pro	3.4
					3.4
		Hs.89925	L04569	calcium channel; voltage-dependent; L ty	
	130568	Hs.16085	AA232535	ESTs; Highly similar to (define not ava	3.4
				ESTs	3.4
		Hs.15978	N78773		
	106036	Hs,10653	AA412505	ESTs	3.4
50					3.4
50		Hs.21893	R45698	ESTs	
	112814	Hs,35828	R98192	ESTs	3.4
				ob93c10.s1 NCI_CGAP_GCB1 Homo sapiens cD	3.4
		Hs.255015			
	100144	Hs.75616	D13643	KIAA0018 gene product	3.4
					3.4
	101129	Hs.247992	L10405	Homo saplens DNA binding protein for sur	
55	130874	Hs.20621	T06287	ESTs	3.4
22					
	106882		AA489009	ESTs	3.4
	103855	Hs.302267	AA195179	ESTs	3.4
		110.0002.01			
	125957		H45213	yo03b08.r1 Soares adult brain N2b5HB55Y	3.3
	114048	Hs.146085	W94613	ESTs	3.3
60					
60		Hs.75354	F13702	ESTs	3.3
		Hs.170098	B45630	ESTs: Highly similar to KIAA0372 (H.sapi	3.3
	104182	Hs.143792	AA479990	ESTs; Weakly similar to glioma amplified	3.3
		Hs.75454	D49396	Human mRNA for Apo1_Human (MER5(Acp1-Mou	3.3
	131688	Hs.30692	U24153	p21 (CDKN1A)-activated kinase 2	3.3
65		Hs.88201	AA481256	ESTs; Weakly similar to (defline not ava	3.3
UJ.				Lo . v, comity distinct to (venille not ave	
	102034	Hs.230	U05291	fibromodulin	3.3
		Hs.14658	R99806	Human chromosome 5q13,1 clone 5G8 mRNA	3.3
	114615	Hs.159456	AA083812	ESTs; Highly similar to (defline not ave	3.3
		Hs.104105		Meis (mouse) homolog 2	3.3
	120/0/	118,104100	nn1004/4	more friends, monotoff 5	3.0

	115048	Hs.190057	AA252668	ESTs	3.3
		Hs.31110		ESTs	3.3
	135142	Hs,24192		ESTs	3.3
		Hs.2877	X63629	cadherin 3; P-cadherin (placental)	3.3
5		Hs.62604		ESTs	3.3
	100365	Hs.79284	D78611	mesoderm specific transcript (mouse) horn	3.3
		Hs.301804		ESTs .	3.3
		Hs.159627		Death associated protein 3	3.3
		Hs.182575		solute carrier family 15 (H+/peptide tra	3.3
10		Hs.140942		ESTs	3.3
			HG3264-HT3441		3.3
				Homo sapiens BAC clone RG041D11 from 7q2	3.3
	102927	Hs.65114	X128/6	keratin 18	3.3
15	132616	Hs.129781		ESTs ESTs	3.3
13		Hs.31652		ESTs	3.3
			AA243681	ESTs	3.3
		Hs.112227		ESTs	3.3
		Hs.12315		ESTs	3.3
20		Hs.178604		ESTs	3.3
20			AA169640	Homo sapiens mRNA for KIAA0643 protein;	3.3
	107391	Hs.284294	W02877	ESTs	3.3
	113058	Hs.7569	T26893	EST	3.3
	134371	Hs.82318	S69790	Brush-1	3.3
25	125669	Hs.333256	R51308	ESTs; Moderately similar to !!!! ALU SUB	3.3
	111506	Hs.294105	R07726	ESTs	3.3
	122974	Hs.194215	AA478625	ESTs	3.3
	102369	Hs.299867	U39840	hepatocyte nuclear factor 3; alpha	3.3
				ESTs	3.3
30		Hs.47402		ESTs; Weakly similar to IIII ALU SUBFAMI	3.3
		Hs.11500	AA437118	ESTs	3.3
		Hs.126494		ESTs	3.3
	127265		AA332751	EST37214 Embryo, 8 week I Homo sapiens c	3.3
25		Hs.41143		Homo sapiens mRNA for KIAA0581 protein;	3.2
35	104856	Hs.293691		ESTs	3.2
		Hs.250655	A97303 AA458761	H.sapiens mRNA for Ptg-12 protein ESTs	3.2
		Hs.251946		ESTs	3.2
	100010	Hs.44481	1112220	forkhead (Drosophila)-like 6	3.2
40	102010	Hs.32425	V V V V V V V V V V V V V V V V V V V	ESTs	3.2
40		Hs.169780		homologous to yeast nitrogen permease (c	3.2
		Hs.292581		ESTs	3.2
		Hs.264207		ESTs	3.2
		Hs.105116		EST	3.2
45			AA598745	ESTs	3.2
		Hs.194657		H.sapiens gene encoding E-cadherin, exon	3.2
		Hs.270016		ESTs	3.2
		Hs.74137	L40391	Homo saplens (clone s153) mRNA fragment	3.2
		Hs.795	L19779	H2A histone family; member O	3.2
50	125596			yg44h11.r2 Soares Infant brain 1NIB Homo	3.2
	127261		AA661567	nu86b02.s1 NCI_CGAP_Alv1 Homo sapiens cD	3.2
		Hs.59554		ESTs	3.2
		Hs.166982			- 32
			AA382283	ESTs	3.2
55		Hs.274256		ESTs	3.2 3.2
		Hs.191185 Hs.99913		ESTs adrenergio: beta-1-; receptor	3.2
		Hs.278834	J03019	Human mDNA for KIA 50146 cone: cortial of	3.2
		Hs.192803	D14592	Human mRNA for KIAA0146 gene; partial cd xeroderma pigmentosum; complementation g	3.2
60		Hs.84072	1147792	transmembrane 4 superfamily member 3	3.2
50		Hs. 116774		integrin; alpha 1	3.2
		Hs.24095		ESTs	3.2
				H3 histone family; member K	3.2
		Hs.189716	R11499	ESTs	3.2
65	121515	Hs. 104696	AA412133	ESTs	3.5
	107445	Hs.6639	W28406	ESTs	3.1
	106887	Hs.334335	AA489091	ESTs	3.2
			AA481806	ESTs	3.
	107072	Hs.130760	AA609113	Homo sapiens mRNA; cDNA DKFZp586N0318 (f	3.2

	102214	Hs.32964	U23752	SRY (sex-determining region Y)-box 11	3.2
	123147		AA487961	ab11h6.s1 Stratagene lung (#93721) Homo	3.2
		Hs.272138		ye87g03.r1 Scares fetal liver spleen 1NF	3.2
_		Hs.250646		ESTs; Highly similar to ubiquitin-conjug	3.2
5		Hs.180789		Homo sapiens (clone S164) mRNA; 3' end o	3.2
		Hs,78344		myosin; heavy polypeptide 11; smooth mus	3.2
		Hs.304389		ESTs Homo sapiens mRNA; chromosome 1 specific	3.2
		Hs.67619	AA069559	colony stimulating factor 1 (macrophage)	3.2
10		Hs.182378 Hs.242894		ADP-ribosylation factor-like 1	3.1
10		Hs.234896		ESTs; Highly similar to geminin (H.sapie	3.1
		Hs.5669	C14290	ESTs	3.1
		Hs.227933		ESTs; Highly similar to (defilne not ava	3.1
		Hs.239720		ESTs; Weakly similar to Rga [D.melanogas	3.1
15		Hs.16346	AA234100	ESTs	3.1
	100631	Hs.48332		Serine/Threonine Kinase (Gb:Z25431)	3.1
	130791	Hs.199263	AA259102	ESTs; Highly similar to (defline not ava	3.1
		Hs.300855		ESTs	3.1
		Hs.123642		ESTs	3.1
20		Hs.98968	Al494372	ESTs	3.1
		Hs.79136	U41060	Human breast cancer; estrogen regulated	3.1
		Hs.47334	W85888	ESTs; Moderately similar to IIII ALU SUB	3.1
		Hs.296842		ESTs; Moderately similar to non-muscle m	3.1
25	125863	Hs.40719 Hs.288192	AA299096	Homo sapiens mRNA; cDNA DKFZp564M0916 (f ESTs	3.1
23		Hs.296141		ESTS	3.1
		Hs.178294		ESTs	3.1
	107332	Hs.183297	T87750	ESTs	3.1
	123570	Hs.109653	AA608955	ESTs	3.1
30		Hs.90800		matrix metalloproteinase 16 (membrane-in	3.1
		Hs.38972	AA161043	tetraspan 1	3.1
	133284	Hs.182828	U09367	zinc finger protein 136 (clone pHZ-20)	3.1
		Hs.33010	H80622	Homo sapiens mRNA for KIAA0833 protein;	3.1
~ -	117606	Hs.44698	N35115	ESTs	3.1
35		Hs.287849		ESTs	3.1
		Hs.103120		ESTs	3.1 3.1
	100789	11- 450440		Phosphoglucomutase 1, Alt. Splice	3.1
	126017	Hs.159440 Hs.247324	M60487	ESTs Homo saplens DNA sequence from PAC 262D1	3.1
40		Hs.108479		ESTs	3.1
40		Hs.181368		U5 snRNP-specific protein (220 kD); orth	3.1
		Hs.118258		ESTs	3.1
	123465	1101110000	AA599033	ESTs	3.1
	126486	Hs.152316	AA345339	EST51345 Gall bladder II Homo sapiens cD	3.1
45	126480	Hs.167031	W01616	za36d05.r1 Soares fetal liver spieen 1NF	3.1
		Hs.43234	N72094	ESTs	3.1
	103860	Hs.38057	AA203742	ESTs	3.1
		Hs.124347		ESTs	3.1
50		Hs.223241		yb15c11.s1 Stratagene placenta (#937225)	3.1
50		Hs.15220		j312.seq.F Human fetal heart, Lambda ZAP	3.1 3.1
		Hs.22242	AA463737 AA442604	ESTs ESTs; Weakly similar to Ydr374cp [S.cere	3.1
	114032	Hs.20993 Hs.35014	W92779	ESTs Vielany Stillian to Full 74cp (5.0016	- 3
	128835	Hs.106390		ESTs	3
55	103667			H.sapiens H4/I gene	3
50		Hs.250614		yy13h06.r1 Soares melanocyte 2NbHM Homo	3
		Hs.21275	D25755	ESTs	3
		Hs.75354	N87590	ESTs	3
		Hs.5811	R12421	ESTs	3
60		Hs.22116	AA307744	Homo sapiens Cdc14B1 phosphatase mRNA; c	3
		Hs.84063	AA016186	ESTs	3
		Hs.8867	U62015	Homo sapiens Cyr61 mRNA, complete cds	3
		Hs24336	W37999	ESTs	3
65		Hs.301404		RNA binding motif protein 3 ESTs	3
65		Hs.146170 Hs.79411	J05249	replication protein A2 (32kD)	3
		Hs.79411 Hs.248177		Human histone H3 gene	3
		Hs.30738	AA257971	ESTs	3
		Hs.33287	U85193	nuclear factor VB	3

		Hs.241551		ESTs	3
		Hs.24104	R26708	ESTs granzyme K (serine protease; granzyme 3;	3
		Hs.3066 Hs.21291	U26174 HG2706-HT2802		3
5		Hs.58915	W86838	EST (GENELES-LES)	3
		Hs.118281		zinc finger protein 288	3
	133780	Hs.76152	M14219	decorin	3
		Hs.14449	AA010889	ESTs	3
		Hs.304139		EST	3
10		Hs.116346		ESTs	3
		Hs.143880		ESTs ESTs	3
		Hs.187555 Hs.50748	T71561	ESTS	3
		Hs.103804		heterogeneous nuclear ribonucleoprotein	3
15		Hs.251531		proteasome (prosome; macropain) subunit;	3
		Hs.10450	AA621125	Homo sapiens chromosome 2; 10 repeat reg	3
		Hs.22545	R43910	ESTs	3
		Hs.263727		ESTs; Moderately similar to !!!! ALU SUB	3
20		Hs.21739		Homo sapiens mRNA; cDNA DKFZp58611518 (f	3
20	131230		AA149987 N79435	thymus specific serine peptidase ESTs	3
	133743	Hs.75847 Hs.227949		ESTs; Highly similar to SEC13-RELATED PR	3
		Hs.44189	N30426	ESTs	3
		Hs.112699		ESTs	3
25		Hs.63290	AA298588	EST114219 HSC172 cells II Homo sapiens c	3
	103795		AA112222	ESTs; Moderately similar to (define not	3
	115092		AA255903	CD39-like 4	2.9
	134831		S72370	pyruvate carboxylase	2.9
30	128579	Hs.101810 Hs.7980	F09570	ESTs; Weakly similar to !!!! ALU SUBFAMI ESTs	2.9
30	123522			ESTs	2.9
		Hs.32793	AA609943	ESTs	2.9
		Hs.88556	D50405	histone deacetylase 1	2.9
	134399	Hs.82689	H99801	tumor rejection antigen (gp96) 1	2.9
35		Hs.174139		H. saplens RNA for CLCN3	2.9
		Hs.14512		ESTs (SCOTES)	2.9
	108555	Hs.2110	AA084963 HG945-HT945	zn13e12.s1 Stratagene hNT neuron (#93723 Nucleic Acid-Binding Protein (Gb:L12993)	2.9
		Hs.16492	AA173998	ESTs; Weakly similar to weakly similar t	2.9
40		Hs.139226		replication factor C (activator 1) 2 (40	2.9
	106636		AA459950	ESTs	2.9
	129109	Hs.108708	AA491295	calcium/calmodulin-dependent protein kin	2.9
		Hs.251871		stromal cell-derived factor 1	2.9
45		Hs.9857	AA433946	ESTs; Weakly similar to (deffine not ava	2.9
45	100386			peroxisomal biogenesis factor 6 ESTs; Moderately similar to !!!! ALU SUB	2.9
	114546		AA056263 AA402224	Homo sapiens growth arrest and DNA-damag	2.9
	108552	ns.3701	AA084912	zn11c7.s1 Stratagene hNT neuron (#937233	2.9
		Hs.190057		16a11 Human retina cDNA randomly primed	2.9
50	134098	Hs.79086	X06323	Human MRL3 mRNA for ribosomal protein L3	2.9
	129721			eukaryotic translation initiation factor	2.9
		Hs.277422		Homo saplens mRNA for cadherin FIB3, par	2.9
		Hs.44104	N29862	ESTs : Moderately similar to WAP four-dis	2.9
55	106335	Hs.36688 Hs.250870	AA437258	protein kinase; mitogen-activated; kinas	2.9
55	105835		AA398412	ESTs	2.9
	106611		AA458904	ESTs; Weakly similar to torsinA (H.sapie	2.9
		Hs.173824		thymine-DNA glycosylase	2.9
	100641	Hs.182183		Caldesmon 1, Alt. Splice 4, Non-Muscle	2.9
60	104602		R86920	ESTs	2.9
		Hs.42738	H99799	ESTs	2.9
		Hs.34073 Hs.155212	AA401912 M65131	BH-protocadherin (brain-heart) methylmalonyl Coenzyme A mutase	2.9
		Hs.5724	AA279422	ESTs	2.9
65	125812			lactin; mannose-binding; 1	2.9
50			H99675	ESTs	2.9
		Hs.285728		H.sapiens mRNA for ArgBPIB protein	2.9
		Hs.132390		ESTs	2.9
	102772	Hs.161002	U63115	absent in melanoma 1	2.9

	131710	Hs.30985	AA233225	ESTs; Highly similar to (defline not ava	2.9
	125231	Hs.268903		ESTs	2.9
		Hs.15535	AI417137	Homo sapiens clone 24582 mRNA sequence	2.9
~		Hs.61289	AB002346	inositol phosphate 5'-phosphatase 2 (syn	2.9
5		Hs.191385		ESTs	2.9
		Hs.303030		EST	2.9
		Hs.34578	AA187045	ESTs; Weakly similar to IIII ALU SUBFAMI	2.9
		Hs.78961	U14575	protein phosphatase 1; regulatory (inhib	2.9
10		Hs.107815		ESTs ESTs	2.9
10		Hs.303125 Hs.218329		heet shock 70kD protein 1	2.9
		Hs.75462		Human BTG2 (BTG2) mRNA; complete cds	2.9
		Hs.18271	AA191014	ESTs; Weekly similar to Ydr372cp [S.cere	2.9
		Hs.232068		Human mRNA for transcription factor AREB	2.9
15		Hs.336901		ESTs	2.9
		Hs.37637	N59645	ESTs	2.9
		Hs.11805	N66066	ESTs	2.9
	128639	Hs.102897	N91246	ESTs	2.9
		Hs.79265	AA114183	ESTs; Moderately similar to glutamate py	2.9
20	135154	Hs.267812	AA126433	sorting nexin 4	2.9
		Hs.279609		pigment epithelium-derived factor	2.9
		Hs.106149		ESTs	2.9
		Hs.2128	U15932	dual specificity phosphatase 5	2.9
05	128104		AA971000	op67g11.s1 Soares_NFL_T_GBC_S1 Homo sapi	2.8
25		Hs.337631		nz22c08.s1 NCI_CGAP_GC81 Homo sapiens cD	2.8
		Hs.180952		ESTs	2.8
		Hs.217916 Hs.93883	D10537	ESTs myelin protein zero (Charcot-Marie-Toolh	2.8
		Hs.68644	N45014	yy60g06.r1 Soares_multiple_sclerosis_2Nb	2.8
30	121873		AA426270	ESTs	2.8
50		Hs.98684	AA432141	ESTs	2.8
		Hs.322645		ESTs	2.8
	135400	Hs.99915	M23263	androgen receptor (dihydrotestosterone r	2.8
		Hs.129998		ESTs	2.8
35		Hs.109019		ESTs	2.8
		Hs.12186	R45480	cyclin K	2.8
	H45968	Hs.32149	H45968	ESTs	2.8
	104261		AF008442	RNA polymerase I subunit	2.8
		Hs.282093		ESTs	2.8
40		Hs.5957	AA417761	Homo saplens clone 24416 mRNA sequence	2.8
		Hs.25960	M13241	v-myc avian myelocytomatosis viral relat	2.8
		Hs.26255	R42714	EST	2.8
	133199		AA609773	Homo sapiens clone 23904 mRNA sequence	2.8
45		Hs.33130	H44825 AA236843	ESTs: Weakly similar to unknown [S.cerev	2.8
43		Hs.72065	R20353	yg20f10.r1 Soares infant brain 1NiB Homo	2.8
	128152	Hs.23740	AA598710	ESTs	2.8
		Hs.97101	AA215333	ESTS	2.8
		Hs.184510		stratifin	2.8
50	132020		AA428990	ESTs	2.8
-	116354	Hs.292566	AA504262	ESTs	2.8
	125867		H98141	ESTs	2.8
	120603	Hs.98541	AA282787	ESTs; Highly similar to (defline not ava -	2.8
		Hs.46847	AA256524	Human DNA sequence from clone 30M3 on ch	2.8
55		Hs.170290		discs; large (Drosophila) homolog 5	2.8
		Hs.110826		Homo sapiens CAGF9 mRNA; partial cds	2.8
	128687		Z38910	ESTs	2.8
		Hs.10299	H09594	ESTs; Moderately similar to III ALU SUB	2.8
		Hs.66731	U81599	homeo box B13	2.8
60		Hs.336829		ESTs; Weakly similar to zinc finger prot FST	2.8
		Hs.25067			2.8
		Hs.173694 Hs.6019	AA430108	ESTs; Moderately similar to (defline not ESTs	2.8
		Hs.22564	AA160890	myosin VI	2.8
65		Hs.40919	N94527	ESTs	2.8
05		Hs.1594	U14518	centromere protein A (17kD)	2.8
		Hs.79981	U79242	Human clone 23560 mRNA sequence	2.8
	129387	Hs.274324		PCAF associated factor 65 alpha	2.8
	126663	Hs.181297	AA714635	ESTs	2.8

	104367	Hs.134342	H17438	ESTs; Weakly similar to seventransmembra	2.8
		Hs.193700		ESTs; Moderately similar to III! ALU SUB	2.8
		Hs.145096		ESTs	2.8
	124447		N48000	ESTs	2.8
5		Hs.125565		deafness; X-linked 1; progressive	2.8
-		Hs.79018		chromatin assembly factor I (150 kDa)	2.8
		Hs.100912		ESTs	2.8
		Hs.326416		ESTs	2.8
			AA402482	ESTs	2.8
10		Hs.75319	X59618	ribonuclectide reductase M2 polypeptide	2.8
10		Hs.35198		ESTs	2.8
		Hs.35380	H88496	ESTs ·	2.8
		Hs.62245		solute carrier family 25 (mitochondrial	2.8
	104000	Hs.29669	AA004000	ESTs	2.8
15		Hs.97694		ESTs	2.8
13		Hs.243901		ESTs	2.8
		Hs.22869		ESTS	2.8
		Hs.168818		ESTs; Moderately similar to roundabout 1	2.8
					2.8
20		Hs.181444 Hs.190478		ESTs; Weakly similar to R12C12.6 [C.eleg ESTs	2.8
20				collagen; type IV; alpha 3 (Goodpasture	2.8
	132598		M81379		2.8
		Hs.1313	L09753	tumor necrosis factor (ligand) superfami ESTs	2.8
		Hs.105640			2.6
25	121329	Hs.1755	AA404324	ESTs Control D	2.7
25	100481	HS.121489	HG1098-HT1098	Cystein D	2.7
		Hs.283683		ESTs	2.7
		Hs,169001		ESTs; Weakly similar to cytochrome P-450	2.7
	432888		T86823	ESTs	2.7
20		Hs.188898		ESTs	2.7
30		Hs.155313		Human mRNA for KIAA0333 gene; partial cd	
			AA398936	ESTs; Weakly similar to (defline not ava	2.7
	131129		R27296	ESTs	2.7
		Hs.272429		calcium-sensing receptor (hypocalcium	2.7
25		Hs.87819		ESTs; Weakly similar to keratin 9; cytos	2.7
35	111900	Hs.25318	R39044	ESTs	
	106025	Hs.173334	AA412063	ESTs	2.7
		Hs.40639		yx92a07.r1 Soares melanocyte 2NbHM Homo	2.7
		Hs.75262	X77383	cathepsin O	2.7
40		Hs.274170		Homo saplens Opa-interacting protein OIP	
40	101584	Hs.84072	M35252	transmembrane 4 superfamily member 3	2.7
		Hs.167489		ESTs	2.7
			AA130156	ESTs	2.7
		Hs.9973	W92797	ESTs	2.7
		Hs.132967		ESTs	2.7
45	134579	Hs.85963	N23222	ESTs; Moderately similar to IIII ALU SUB	2.7
		Hs.258301		ESTs	2.7
		Hs.332541		ESTs; Weakly similar to HEM45 [H.sapiens	2.7
		Hs.179825		Human sperm membrane protein BS-63 mRNA,	2.7
~~		Hs.99598	AA463627	ESTs	2.7
50	134983	Hs.196384	D26235	prostaglandin-endoperoxide synthase 2 (p	2.7
	120537	Hs.160422	AA262790	ESTs	2.7
	131036	Hs.174140	X64330	ATP citrate lyase	2.7
	133889	Hs.211582	AA099391	ESTs	2.7
		Hs.106529		zv81e01.r1 Soares_total_felus_Nb2HF8_9w	2.7
55		Hs.306044		ESTs	2.7
	423239		AA323591	EST26392 Cerebellum II Homo seplens cDNA	2.7
	105031	Hs.12321	AA127240	ESTs	2.7
	126021	Hs.187516		ESTs	2.7
	102116		U13706	Human ELAV-like neuronal protein 1 isofo	2.7
60		Hs.237225		ESTs; Weakly similar to (defline not ava	2.7
		Hs.278439		ESTs	2.7
		Hs.40241	AA004878	ESTs; Highly similar to (defline not ava	2.7
		Hs.1259	X55283	asialoglycoprotein receptor 2	2.7
	112109	Hs.283309		ESTs: Weakly similar to filf ALU SUBFAMI	2.7
65	128422		T85681	yd60c06.r1 Soares fetal liver spleen 1NF	2.7
	109494	Hs.43899	AA233702	ESTs	2.7
	118696	Hs.292284	N72086	Homo sapiens RNA polymerase III largest	2.7
		Hs.36727	AA416963	ESTs; Highly similar to histone H2A [H.s	2.7
		Hs.284380	L20492	gamma-glutamyltransferase 1	2.7

		Hs.111323		EST; Highly similar to (defline not avai	2.7
	123798		AA620411	smell inducible cytokine A5 (RANTES)	2.7
	106716	Hs.238928		ESTs	2.7
-	103663		Z78291	Z78291 Homo saplens brain fetus Homo sap	2.7
5		Hs.22265	Z38909	ESTs	2.7 2.7
		Hs.5027	T32438	ESTs af60c09.r1 Soares_NhHMPu_S1 Homo sapiens	2.7
	127897 130621	Hs.16803	AA773857 AA621718	ESTs: Weakly similar to (define not ava	2.7
		Hs.42796	AA479958	ESTs; Highly similar to (define not ava	2.7
10	125499	118.46700	R11878	yf49d11.r1 Soares infant brain 1NIB Homo	2.7
10		Hs.77899	M19267	tropomyosin 1 (alpha)	2.7
	104470			ESTs; Weakly similar to Similar to colla	2.7
	134982		N48088	ESTs	2.7
		Hs.284295		ESTs	2.7
15		Hs.285574		ESTs	2.7
	125401	Hs.337585	Al204637	ESTs; Moderately similar to KIAA0350 [H.	2.7
		Hs.15768	N70042	ESTs; Moderately similar to !!!! ALU SUB	2.7
		Hs.164478		ESTs; Weakly similar to (defline not ava	2.7
	134507		M63488	replication protein A1 (70kD)	2.7
20	121609	Hs.98185	AA416867	EST	2.7
	113835	Hs.27475	W56590	ESTs	2.7
		Hs.285290	AA428062	ESTs; Highly similar to (defline not ava	2.7 2.7
		Hs.98558 Hs.216717		ESTs ESTs	2.7
25		Hs.12698	AA464273	ESTS	2.7
23	123184		AA489072	Homo saplens mRNA for KIAA0870 protein;	2.7
		Hs.173497		SEC23-like protein B	2.7
	106186		AA427398	acetylserotonin N-methyltransferase-like	2.7
	101349	11010010	L77559	Homo sapiens DGS-B partial mRNA	2.7
30	112954	Hs.6655	T16559	ESTs	2.7
	133054	Hs.291079	R07876	ESTs; Weakly similar to unknown [S.cerev	2.7
	128131	Hs.25640	Al283162	claudin 3	2.6
	101864		M95787	transgelin	2.6
		Hs.26303	R40752	ESTs	2.6
35		Hs.151051		protein kinase mitogen-activated 10 (MAP	2.6
		Hs.23964	Al362218	ESTs	2.6
		Hs.47111	N50740	ESTs	2.6 2.6
	116345 132227	Hs.199067 Hs.4248	AA496981 AA412620	ESTs ESTs	2.6
40	132227			yj42b06.r1 Scares placenta Nb2HP Homo sa	2.6
70		Hs.89463	AA137034	ESTs	2.6
	102764	110,00400	U82310	Homo saplens unknown protein mRNA, parti	2.6
		Hs.173933		ESTs	2.6
			AA307896	nuclear localization signal deleted in v	2.6
45	107427	Hs.46736	W26975	ESTs	2.6
	117477	Hs.44175	N30328	ESTs	2.6
	103290	Hs.16364	AA435542	ESTs	2.6
	126829		R11547	ESTs	2.6
~~	118836			ESTs	2.6
50		Hs.136348		osteoblast specific factor 2 (fasciclin	2.6
	104278			ESTs; Highly similar to (defline not ava	2.6 2.6
	135051		C15324	ESTs collagen: type I: alpha 1	- 2.6
	123579	⊓8.227 <b>63</b> 3	AA608983	af5d4.s1 Soares_testis_NHT Homo sapiens	2.6
55		Hs.149923		X-box binding protein 1	2.6
33	101434		M20218	coagulation factor XI (plasma thrombopla	2.6
	122962			ESTs; Moderately similar to !!!! ALU SUB	2.6
	126151		AA324743	ESTs	2.6
	128925	Hs.21851	D61676	Homo saplens mRNA; cDNA DKFZp586.i2118 (f	2.6
60		Hs.103391		Insulin-like growth factor binding prote	2.6
		Hs.154103		LIM protein (similar to rat protein kina	2.6
	128402			ESTs	2.6
	129273			ESTs	2.6
ce	125483		F07759	ESTs	2.6
65	132953		AA029927	ESTs	2.6
	130963		U57099	nuclear protein; marker for differentiat	2.6 2.6
	120614	Hs.194154 Hs.103267	AA284281 AA490988	ESTs; Weakly similar to IIII ALU SUBFAMI ESTs; Moderately similar to Rabin3 (R.no	2.6
		Hs.96744		ESTS	2.6
	.21710				

	125428	Hs.851	W74608	ESTs; Highly similar to (defline not ava	2.6
	115906	Hs.82302	AA438616	ESTs	2.6
	103432		AA076626	Homo sapiens clone 23851 mRNA sequence ESTs	2.6 2.6
5		Hs.191911 Hs.281434		ESTs	2.6
5		Hs.268615		ESTs	2.6
		Hs.173840		ESTs	2.6
	102565		U59748	Human desert hedgehog (hDHH) mRNA, parti	2.6
10		Hs.13109	AA194973	ESTs	2.6
10	114264	Hs.334609 Hs.21104	AA429951	ESTs ESTs	2.6 2.6
	135192			purinergic receptor P2X; ligand-gated to	2.6
	109833			ESTs	2.6
		Hs.8535	AA303088	ESTs; Weakly similar to transformation-r	2.6
15		Hs.97967	AA406210	ESTs	2.6
		Hs.155485		Human huntingtin interacting protein (HI	2.6
		Hs.102329 Hs.97199		ESTs ESTs	2.6
	127081		RRRSR2	ESTs; Weakly similar to weak similarity	2.6
20		Hs.11463	AA458603	ESTs; Weakly similar to (defline not ava.	2.6
	112410	Hs.26904	R61680	ESTs	2.6
		Hs.112981		ESTs	2.6
	122905	Hs.104835	AA470070	ESTs	2.6 2.6
25	116399	Hs.110637 Hs.153934	AA599729	Homo saplens homeobox protein A10 (HOXA1 core-bloding factor; runt domain; alpha	2,6
23		Hs.1435	M24470	quanosine monophosphate reductase	2.6
				Trithorax Homolog Hrx	2.6
	104965		AA084104	ESTs	2.6
	117711		N45201	EST	2,6
30	124792		R44357 N73808	ESTs ESTs	2.6 2.6
	111299	Hs.74313 Hs.32971	Z46973	phosphoinositide-3-kinase; class 3	2.6
	133629			KIAA0017 gene product	2.6
		Hs.169977	Al086782	ESTs	2.6
35	100858			Forkhead Family Afx1	2.6
		Hs.301927		T-cell receptor; alpha (V;D;J;C) ESTs	2.6 2.6
		Hs.133865 Hs.92137	AA428557	v-myc avian myelocytomatosis viral oncog	2.6
		Hs.10247	U30999	Human (memc) mRNA, 3'UTR	2.6
40		Hs.191538		ESTs	2.6
		Hs.34136	AA307443	ESTs	2.6
		Hs.268601		ESTs; Weakly similar to (define not ava	2.6
		Hs.21201 Hs.40022	Z39338 H79310	ESTs; Highly similar to (defline not ava. EST	2.6
45		Hs.306995		ESTs	2.6
15	133989	Hs.78202	U29175	SWVSNF related; matrix associated; acti	2.6
				Caldesmon 1, Alt. Splice 3, Non-Muscle	2.6
		Hs.285996		ESTs	2.6
50		Hs.6540 Hs.171391	Z40861	ESTs C-terminal binding protein 2	2.6
50			AA017258	EST	2.5
	100134		D13264	macrophage scavenger receptor 1	2.5
	133969		U13044	GA-binding protein transcription factor;	- 2.5
		Hs.74316	AA455001	ESTs	2.5
55	127493	Hs.291701	AA808081	oc39a08.s1 NCI_CGAP_GCB1 Homo sapiens cD	2.5
	132869	Hs.203961 Hs.44583	N26855 N34415	ESTs EST	2.5
	124844	Hs.109654		ESTs	2.5
		Hs.2785	Z19574	keratin 17	2.5
60		Hs.5897	AA047151	ESTs	2.5
		Hs.82643	U02680	protein tyrosine kinase 9	2.5
		Hs.20159 Hs.193784	AA454156	ESTs ESTs	2.5
		Hs.193784 Hs.24908	AA430044 AA256042	ESTS	2.5
65		Hs.75275	D50916	homolog of yeast (S. cerevisiae) uld2	2.5
	102959	Hs.121524	X15722	glutathione reductase	2.5
		Hs.6166	AA047616	ESTs	2.5
		Hs.2057 Hs.118131	AA128100	uridine monophosphate synthetase (orotat 5;10-methenyltetrahydrofolate synthetase	2.5
	129045	16.110131	L00920	o, ro-menenyissanyuroromo symbolissa	2.5

	126399	Hs.83883	AA128075	zi16d08.r1 Scares pregnant_uterus_NbHPU	2.5
	134069	Hs.78935	U29607	Homo sapiens elF-2-associated p67 homolo	2.5
	109816	Hs.61960	F11013	ESTs: Weakly similar to KIAA0176 [H.sapl	2.5
	134801	Hs.89695	X02160	insufin receptor	2.5
5	104232	Hs.10587	AB002351	Human mRNA for KIAA0353 gene; partial od	2.5
-	107361	Hs.159486	U72513	Human RPL13-2 pseudogene mRNA; complete	2.5
	106057	Hs.289074	AA417067	ESTs	2.5
	134252	Hs.80720	AA031782	Homo sapiens mRNA; cDNA DKFZp586B1722 (f	2.5
	128062	Hs.105547	AA379500	ESTs	2.5
10	110009	Hs.6614	H10933	ESTs	2.5
	111375	Hs.20432	N93696	ESTs	2.5
	122642	Hs.99361	AA454186	ESTs	2.5
	127999	Hs.69851	AA837495	ESTs: Weakly similar to Wiskott-Aldrich	2.5
	105029	Hs 13268	AA126855	ESTs	2.5
15	105082	Hs.26765	AA143763	ESTs; Weakly similar to Similarity to S.	2.5

TABLE 1A show the accession numbers for those primekeys lacking unigeneID's for Table

1. For each probeset we have listed the gene cluster number from which the oligonucleotides

5 were designed. Gene clusters were compiled using sequences derived from Genbank EST's
and mRNAs. These sequences were clustered based on sequence similarity using Clustering
and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers
for sequences comprising each cluster are listed in the "Accession" column.

10

	Pkey:		Unique Eos probeset Identifier number					
	CAT number:		ene diuster number					
	Accession:		Senbank accession numbers					
15	Pkey	CAT number	Accessions					
	108550	111555 1	AAD71210 AA069699 AAD71438 AA084912 AA084803 AAD79371 AAD79370					
		1596090 1	H57661 H58881					
20		1606216 1	H75681 H70975					
20		32479 1	AB010994 U59748 AA064660					
		48158 -7	S81578					
		1562851 1	H10543 R11878					
		1708455_1	R25698 R56582 R56018					
25		37186_1	AF080229 AF080231 AF080230 AF080232 AF080233 AF080234 BESGC83 AR68743 AW614951 BE467547 AR60833 AR63841 ROSS980 BE7650 BE7650 BE7650 CHORD SEARCH URFSS AAR63862 AW608023 AR703576 AR6375 AR602562 AR63864 AW608020 AR703576 AR6375 AR602562 AR63864 AR602564 AW608602 AR602564 AW609564 AR602564 AW609564 AR602564 AR602564 AW609564 AR602564 A					
30			Al352545 BE501030 Al652535 BE465762 AA206331 AW451866 AA471088 AA206342 AA204834 AA206100 AW021661 AA332922 N66048 AA703396 H92278 AW139734 H92683 U87589 U87595 H69001 U87594 BE466420 Al624817					
-			BF488611 AIZ06344 AA574397 AA348354 A493192					
		327827_1 1583542 1	AA491830 R50173 R55192 R50320 Al732306 Al732305 Al820727 Al820728 R55191 R50319 R50227 H41694 H45213					
	125982	1766315_1	R98091 W92898					
35	127248	227560 1	AA364195 AA325029 AW962050					
	103731	112052_1	AA070545 AA131490 AA131373					
	127261	231687_1	AA330501 AA661567					
	127265	232391_1	AA331503 AA332751 AW962542					
	126659	1541209_1	T16245 R19694 F13545 H10299 T66048 T65279 H18006					
40	127315	37938 1	AF116622 Al114507 AA640834 AA377999					
	103806	112618 1	AA130614 AA071410					
	128104	502608 1	AA906093 AA971000					
		524482 2	H47610 R86920					
	128152	297868 1	F07973 R20353 AA442660					
45		1811283 1	T77794 T85881					
		446527_1	AA773681 AA773857					
		120358_1	BE298210 AIG72315 AW086489 BE298417 AA455921 AA602537 BE327124 R14963 AA095210 AW274273 AI333584 AI369742 AI039658 AI885095 AI476470 AI287650 AI885299 AI985381 AW592624 AW340136 AI266556 AA456390					
			Al310615 AA484951					
50	129735	44573_2	AGRICUST INTODOS ROYOU NORDO AUGOS 110 AMERIOT NESSOO AUGUS 1473 ESSOS AMENTATOS TRATZTO MOTIOSI AMERIOTA I AAGUSSIS ALAGEISTA AIRCOST AIRCOSS WISTON 1 RESSOU ATHAS TRASSO I BECREST I TTS TOR TRATZE AABRICUS BECRISTA TACTISTOS AABRIANA MERIOTA TIANGA ARAZTINA AIRCOSTA AIRCOST A					
55			AA856975 AW505512 AI961530 AW629970 BE612881 AW276997 AW513601 AW512843 AA044209 AW656538					
			AA 180000 AAS37469 AWB1101 AAS1669 AAS51874 A1819228 AWC65962 AB63333 ABB5505 AWC78005 AB533030 A A9472894 AM59814 AWK73626 AW519369 AA328274 AA0896759 H7526 AW2628 H584729 H5805 T20487 AM262656 AA780419 AAS5105 Y80070 AWK7366 H5273032 AI564289 F00551 H63488 W37181 W75802 R66056 AA002639 R6780 AA300207 AWK95661 T58226 F04005					
60	123147	2198022	AA487961					
	130529	158447_1	AA178953 AA192740					
	123579	genbank_AA608	983 AA608983					
	109175	genbank_AA180	496 AA180496					
		tigr HT4163	S67998					
65		tigr_HT4515	U10072					

	123798	579959_1	AA620411 AA287491
		entrez_U13706	U13706
		entrez_U42359	U42359
		entrez_U82310	
5	118475	genbank_N66845	N66B45
		genbank_AA026349	
	104787	genbank_AA027317	AA027317
		genbank_T97307	
		genbank_W81598	
10	122835	genbank_AA454085	AA454085
	108407	genbank_AA075519	AA075519
		genbank_AA076626	
	108555	genbank_AA084963	AA084963
	101349	entrez_L77559	L77559
15	124447	genbank_N48000	N48000
	119071	genbank_R31180	R31180
	103520	entrez_Y10511	Y10511
		genbank_Z78291	
			AA873285 Al025762
20			AA199853 AA206355
	123465	genbank_AA599033	AA599033

## MISSING AT THE TIME OF PUBLICATION

TABLE 2: shows a preferred subset of the Accession numbers for genes found in Table 1 which are differentially expressed in prostate tumor tissue compared to normal prostate tissue.

5

102398 U42359

101201 L22524

101803 M86546

65

Unique Eos probeset identifier number ExAcon: Exemplar Accession number, Gentrank accession number UnigeneID: Unigene number 10 Unigene Title: Unigene gene title Ratio of turnor to normal body tissue (Relaxed ratio (87/70) R1: Pkey ExAcon UnigenelD Unigene Title R1 15 372 131919 AA121266 Hs.272458 ESTs 120328 AA196979 Hs.290905 ESTs: Weakly similar to (defline not ava 326 101486 M24902 Hs.1852 acid phosphatase; prostate Hs.279477 ESTs 25.2 119073 R32894 20 238 133428 M34376 Hs.183752 microseminoprotein; beta-128180 AA595348 Hs.171995 kallikrein 3; (prostate specific antigen 21.4 104080 AA402971 Hs.57771 Homo sapiens mRNA for serine protease (T 18.9 18.6 127537 AA569531 He 162859 ESTs 131665 R22139 Hs.30343 ESTs 174 25 101050 K01911 Hs.1832 neuropeptide Y 17.3 130771 N48056 Hs.1915 folate hydrolase (prostate-specific memb 17 16.7 107485 W63793 Hs.262476 S-adenosylmethionine decarboxylase 1 106155 AA425309 Hs.33287 ESTs 165 Hs.11260 ESTs 129534 R73640 16.4 Anticen, Prostate Specific, Alt. Splice 16 30 100589 HG2261-HT2351 101889 \$39329 Hs.181350 kalikrein 2; prostatic 15.4 135389 U05237 Hs.99872 fetal Alzheimer antigen 133944 AA045870 Hs.7780 ESTs 195 130974 X57985 Hs.2178 H2B histone family: member Q 11.8 35 114768 AA149007 Hs.182339 ESTs 11.8 104660 AA007160 Hs.14846 ESTs 114 Hs.268744 ESTs; Moderately similar to KIAA0273 [H. 131061 N64328 109 126645 Al167942 Hs.61635 Homo saplens BAC clone RG041D11 from 7g2 10.7 135153 N40141 Hs.95420 Homo sapiens mRNA for JM27 protein; comp 10.6 40 Hs.113314 ESTs 106 107033 AA599629 ESTs; Weakly similar to polymerase [H.sa 10.5 118417 N66048 126758 W37145 Hs.293960 ESTs 10.2 Hs.30652 ESTs 107102 AA609723 10.1 116787 H28581 Hs.15641 ESTs 45 115719 AA416997 Hs.59622 ESTs 10 123209 AA489711 Hs.203270 ESTs 101664 M60752 Hs.121017 H2A histone family; member A 9.8 Hs.83883 ESTs 112971 T17185 9.7 117984 N51919 Hs.106778 ESTs 50 129523 M30894 Hs.274509 T-cell receptor; gamma cluster 132964 AA031360 Hs.167133 ESTs 121853 AA425887 Hs.96502 ESTs 119617 W47380 Hs.55999 ESTs 80 105627 AA281245 88 Hs.23317 ESTs Hs.76422 phospholipase A2; group IIA (platelets; 55 101461 M22430 Hs.293185 yz61c5.s1 Scares\_multiple\_sclerosis\_2NbH 124526 N62096 133845 T68510 Hs.76704 ESTs 133354 AA055552 Hs.334762 ESTs; Weakly similar to KIAA0319 [H.sapi 8.1 119018 N95796 Hs.278695 ESTs Hs.66052 CD38 antigen (p45) Hs.23023 ESTs 60 100394 D84276 106579 AA458135 76 114985 AA250737 Hs.72472 ESTs 7.4 112033 R43162 7.1 Hs.22627 ESTs Human N33 protein form 1 (N33) gene, exo

118

Hs.2256 matrix metalloproteinase 7 (matrilysin;

Hs.155691 pre-B-cell leukemia transcription factor

120562 AA280036 Hs.302267 ESTs; Weakly similar to W01A6.c [C.elega

	109112	AA169379	Hs.257924		6.8
		F10707	Hs.326416		6.7
		X07730	Hs.171995	kallikrein 3; (prostate specific antigen	6.6
		AA219134	Hs.26691	ESTs	6.6
5	132902	AA490969	Hs,59838	ESTs	6.6
		U07919	Hs.75746	aldehyde dehydrogenase 6	6.5
		241050		Homo sapiens Mod4p homolog mRNA; comp	let 6.5
		AA010163	Hs.3383	upstream regulatory element binding prot	6.5
	100727	X07290		Human HF.12 gene mRNA	6.3
10		AA421714		Homo sapiens mRNA for KIAA0898 protein;	6.3
	123475	AA599267		ESTs; Weakly similar to ANKYRIN; BRAIN V	6.3
	133061	AB000584	Hs.296638	prostate differentiation factor	6.3
	116429	AA609710	Hs.279923	ESTs; Weakly similar to similar to GTP-b	6.2
		L29008	Hs.878	sorbitol dehydrogenase	6.2
15	104691	AA011176	Hs.37744	ESTs	6.2
	127248	AA325029		EST27953 Cerebellum II Homo sapiens cDN/	16.2
	105500	AA256485	Hs.222399	ESTs	6.1
	130828	AA053400	Hs.203213	ESTs	5.9
	115357	AA281793	Hs.72988	ESTs	5.8
20	116334	AA491457	Hs.48948	ESTs	5.7
	120132	238839	Hs.125019	ESTs; Weakly similar to !!!! ALU SUBFAMI	5.6
	106375	AA443993	Hs.289072	ESTs	5.6
	124777	R41933	Hs.140237	ESTs; Weakly similar to neuronal thread	5.6
	101791	M83822	Hs.62354	Human belge-like protein (BGL) mRNA: par	5.5
25	117698	N41002	Hs.45107	ESTs	5.5
	122041	AA431407	Hs.98732	Homo sapiens Chromosome 16 BAC clone C	IT 5.5
	133723	AA088851	Hs.262476	S-adenosylmethionine decarboxylase 1	5.5
	113938	W81598		ESTs	5.4
	133015	AA047036	Hs.246315	ESTs	5.4
30	108186	AA056482	Hs.7780	ESTs	5.3
		N25110	Hs.326392	Human quanine nucleotide exchange factor	5.3
	104033	AA365031	Hs.98944	ESTs	5.3
	110844	N31952	Hs.167531	ESTs; Weakly similar to (defline not ava	5.3
		H70627		ESTs: Weakly similar to !!!! ALU SUBFAMI	5.3
35	133493	AA284143	Hs.194369	Homo sapiens chromosome 1 atrophin-1 rel	5.3
	129184	W26769		ESTs: Highly similar to (define not ava	5.2
		M21389		keratin 5 (epidermolysis buliosa simplex	5.1
		AA464728		ESTs: Weakly similar to IIII ALU SUBFAMI	5.1
		AA402613	Hs.169119		5.1
40		X91868	Hs.54416	sine oculis homeobox (Drosophila) homolo	5.1
		AA400271		ESTs: Highly similar to (defline not ava	5.1
		AA479362	Hs.47144	ESTs	5
		X07696	Hs.80342	keratin 15	5
		X52541		early growth response 1	5
45		N93839	Hs.39288	ESTs: Weakly similar to !!!! ALU SUBFAMI	5
				,,	

TABLE 2A shows the accession numbers for those primekeys lacking unigeneID's for Table 2. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers for sequences comprising each cluster are listed in the "Accession" column.

	CAT number:		Unique Eos probesel distrillation rumber Gene dissiste marcher Genebank accession numbers	
15	Pkey	CAT number	Accession	
20	118417	37186_1	AF000229 AF000231 AF000200 AF0000202 AF000023 AF000234 BESD(033 AIGS0745 AW614851 BE467547 AI860033 AIGS361 NC90081 U97502 U97503 U97500 U97505 NS4404 U97507 AA85502 AW700002 AIR7070 AIS65716 AIR7574 AI875065 AW705404 AIR50256 AV70120 AI87004 AW70120 AV70120 AA84508 AV70140 AA07575 AA0715 AA67015 AA67015 AA0715 AA67015 AA6	
25	127248 107033	227580_1 235652_1	DE400011 MICROSH MASHASIY MASHASIY MASHASIYA MARISHI AAQSA195 AAQSA195 AAQSA2528 ANWEGGEZO ANWEG	
	102338 113938	entrez_U4235 genbank_W81	9 U42359	

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TABLE 3: shows genes, including expression sequence tags, differentially expressed in prostate tumor tissue compared to normal tissue as analyzed using the Affymetrix/Fos Hu02 GeneChip array. Shown are the relative amounts of each gene expressed in prostate tumor samples and various normal tissue samples showing the highest expression of the gene.

10	Pkey: ExAcon: UnigeneID: Unigene Title: R1:		Unique Eos probosot Identifler numbor Exemplar Accession number, Gerbank accession number Uniquen aumbor Uniquen gena tille Alla Gof Luncer to normal body issue				
15							
	Pkey	ExAcon	UnigenelD	Unigene Title	R1		
20	100235 100570 100819	D12485 D29954 HG2261-HT23 HG4020-HT42	90	phosphodiesterase Vnucleotide pyrophosp KIAA0056 protein Hs.171995 Hs.2387	6.3 5.1 Antigen, Prostate Specific, Alt. Splice Transglutaminase 10.5	9	
25	101247 101416 101447	L00354 L33801 M17254 M21305 M24736	Hs.80247 Hs.78802 Hs.279477 Hs.89546	cholecystokinin glycogen synthase kinase 3 beta v-ets avian erythroblastosis virus E26 o Human alpha satellite and salellite 3 ju selectin E (endothellal adheston molecul	8.5 4.7 4.7 11 9.8		
30	101514 101628 101663 101758	M28214 M57399 M60750 M77836 M81118		RAB3B; member RAS oncogene family pleiotrophin (heparin binding growth fac H2B histone family; member A pyrroline-5-carboxylate reductase 1	6.2 8.4 4.9 5.4 7.5		
35	101817 101888 102031 102052	M88163 M99701 U04898 U07559	Hs.152292 Hs.95243 Hs.2156 Hs.505	SWI/SNF related; matrix associated; acti transcription elongation factor A (SII)- RAR-related orphan receptor A ISL1 transcription factor; LIM/homeodoma	5.5 5.7 13.2 8.9		
40	102233 102302 102348	U24576 U26173 U33052 U37519 U48807	Hs.3844 Hs.79334 Hs.69171 Hs.87539 Hs.2359	LIM domain only 4 nuclear factor; interfeukin 3 regulated protein kinase C-like 2 aldehyde dehydrogenase 8 dual spacificity phosphatase 4	5.8 7.4 8.2 5.9 5.1		
45	102473 102669 102698 102751	U49957 U71207 U75272 U80034 U90914	Hs.180398 Hs.29279 Hs.1867 Hs.68583 Hs.5057	LIM domain-containing preferred transicc eyes absent (Drosophila) homolog 2 progastricsin (pepsinogen C) mitochondrial intermediate peptidase carboxypeptidase D	5.7 9 10.6 15.8 4.9		
50	102869 103031 103043 103093 103376	X02544 X54667 X55733 X60708 X92098 X95240	Hs.572 Hs.123114 Hs.93379 Hs.44926 Hs.323378 Hs.54431	orosomucoid 1	22.6 4.7 4.9 5.8 7.4		
55	103677 103962 104064	Z46629 Z63806 AA298180 AA410529	Hs.2316 Hs.83243 Hs.30732	SRY (sex-determining region Y)-box 9 (ca H.sapiens mRNA for axonemal dynain beavy ESTs ESTs	5.2 4.9 6 6.4 6.8		
60	104301 104769 104851 104896 104956	AF006265 D45332 AA025887 AA040882 AA054228 AA074880	Hs.10290 Hs.23165 Hs.20509	estropen receptor-binding fragment-assoc ESTs; Weakly similar to IIII ALU SUBFAMI US anRNP-specific 40 kDa protein (hPrp8- ESTs; Weakly similar to hypothetical pro	10.5 6.3 4.9 5.8 6.4		
65	104957 104967 105099	AA084506 AA150776 AA233459	Hs.10026 Hs.291000 Hs.23729 Hs.26369	ESTs Homo saplens clone 24405 mRNA sequence	4.8 6.5 7 5.1		

	105304	AA233553	Hs.190325		4.7
	105370	AA236476	Hs.22791		10.3
	105427	AA251330	Hs.28248		5
-	105542	AA261858			8.8
5	105628	AA281251	Hs.79828		5.5 8
		AA281623 AA282138	Hs.6685 Hs.11325		14
	105691	AA287097			6.3
		AA292701	Hs,5364		4.9
10	105808	AA393808		KIAA0438 gene product	7
	105826	AA398243			5
	105903	AA401433			9.9
	105906	AA401633	Hs.22380		11.5
15			Hs.25206	ESTs ESTs	5.1 10.9
15	106094	AA419461 AA425367	Hs.23317 Hs.34892		6.6
	106157	AA426643	Hs.10762		8.5
	106211	AA428240	Hs.126083		8.4
		AA428258	Hs.8769	Homo sapiens mRNA; cDNA DKFZp564E153 (fr	5.7
20	106272	AA432074	Hs.323099	ESTs	5.8
		AA443828	Hs.288856		6.3
	106400		Hs.94109		5.4
	106474	AA450212	Hs.42484	Homo sapiens mRNA; cDNA DKFZp564C053 (fr	9.2 5.6
25	106507 106523	AA452584 AA453441	Hs.31511		4.7
23		AA453628	Hs.37443		4.7
	106557	AA455087	Hs.22247		5.7
	106575	AA458039	Hs.105421		7.2
	106618	AA459249	Hs.8715	ESTs; Weakly similar to Similarity with	5.6
30	106820	AA481037	Hs.12592		5.4
		AA485223	Hs.34892		5.3
		AA505141	Hs.11923	Human DNA sequence from clone 167A19 on	7.5 6.1
		AA609952 AA620504	Hs.12784 Hs.179898		7.1
35		AA621340	Hs.10600		5.2
33	107217	D51095	Hs.35861	DKFZP586E1621 protein	15.1
	107365	U78294		arachidonate 15-lipoxygenase; second typ	4.7
	107630	AA007218	Hs.60178	ESTs	5.3
40		AA016225	Hs.7517	ESTs	4.8
40		AA018042	Hs.252085	Human DNA sequence from clone 141H5 on c	7.6 10.5
	107997 108012	AA037388 AA039616	Hs.82223 Hs.173334		6.5
	108520		Hs.46786	ESTs	7.9
		AA088276	Hs.68826	ESTS	5.6
45		AA100967	Hs.69165	ESTs	6
	108664	AA113349	Hs.69588	EST	6.3
	108677	AA115629	Hs.118531	ESTs	5.9
	108807	AA129968	Hs.49378	ESTs; Weakly similar to PROTEIN PHOSPHAT	5.8 5
50	108910		Hs.337232	ESTs ESTs	12.7
30	108948		Hs.118258		6.8
		AA156790	Hs.262036		15.3
		AA171529	Hs.183887		6.1 ^
	109142	AA176438	Hs.41295	ESTs	5.1
55	109277		Hs.86043	ESTs	5.5
	109342	AA213620	11-407004	Home saplens mRNA; cDNA DKFZp586M1418 (	10.8
		F01811		ESTs; Moderately similar to voltage-gate ESTs	7
	109565 109648		Hs.23648 Hs.7154	ESTS	9.9
60		F10770		Homo sapiens clone 669 unknown mRNA; com	6.4
00	109859		Hs.20792	ESTs	5.3
	110181	H20276	Hs.31742	ESTs	16.8
	110854	N32919	Hs.27931	ESTs	10
	110924	N47938	Hs.12940	yy84a09.s1 Soeres_multiple_solerosis_2Nb	5.6
65	111048		Hs.318584	ESTs Homo sapiens mRNA; cDNA DKFZp434N185 (fr	6.9
	111091		Hs.33032 Hs.99364	ESTs	5
	111164		Hs.122499	ESTs; Weakly similar to !!!! ALU CLASS C	5.6
	111221		Hs.15119	ESTs	6.2

	111348	N90041	Hs.9585	ESTs	5.4
	111353		Hs.6616		5.3
	111495		Hs.9683		5.8
	111540	R08850	Hs.9786		6
5		R10657		KIAA0830 protein	12.
		R10684	Hs.5794	ESTs	7.1 6.2
	111734	R25375 R37460	Hs.128749 Hs.25231	ESTS	9.4
	111870		Hs.18685	ESTs; Weakly similar to hypothetical pro	6.5
10	111937		Hs.14846	Homo sapiens mRNA; cDNA DKFZp564D016 (fr	
	111987	R42036	Hs.6763	KIAA0942 protein	6.4
	112184		Hs.330242	ESTs	5.6
		R53765		KIAA0981 protein	9.3
15		R59740	Hs.5740	ESTs	4.7
15	112452		Hs.157461 Hs.78225		6 5.4
	112753		Hs.169882		5.8
	112902		Hs.129190		5.1
	112984		Hs.289014	ESTs	4.9
20	113021	T23855		KIAA1028 protein	10.
	113083			ESTs; Weakly similar to heat shock prote	5.7
		T57773	Hs.10263 Hs.86538	ESTs ESTs	7.3 8.7
		T88878 W60439	Hs.8858	ESTs; Moderately similar to cbp146 [M.mu	4.9
25		W72382	Hs.11958	oxidative 3 alpha hydroxysteroid dehydro	4.7
		W85765	Hs.30504	Homo saplens mRNA; cDNA DKFZp434E082 (fr	
		W87462	Hs.21894	ESTs	5.9
		W87544	Hs.268828		4.7
30	114124	Z38595 Z41395	Hs.125019 Hs.143611	ESTs; Highly similar to KIAA0886 protein	9.6
50	114346		Hs.130489		5.2
		AA018216		Bicaudal D (Drosophila) homolog 1	7.4
		AA025370	Hs.40109	KIAA0872 protein	8.2
0.5		AA101416	Hs.107149		5.4
35	114721		Hs.103822		4.8
	114730 114833		Hs.331328 Hs.87159	ESTs; Weakly similar to The KIAA0138 gen ESTs; Moderately similar to CGI-66 prote	5.1
	114860		Hs.42179	ESTs; Moderately similar to similar to m	6.3
		AA235811	Hs.293872		5.2
40	114895	AA236177	Hs.76591	KIAA0887 protein	4.7
		AA236545	Hs.54973	ESTs	5.2
		AA242751	Hs.16218 Hs.42484	KIAA0903 protein Homo sapiens mRNA; cDNA DKFZp564C053 (fr	5.7
		AA255566 AA258030		ESTs; Weakly similar to supported by GEN	5.8
45		AA287061	Hs.48499	ESTs; Highly similar to Bdelght protein	4.7
		AA398913	Hs.45231	LDOC1 protein	7.6
		AA412519	Hs.58279	ESTs	4.8
		AA423972	Hs.131740		5
50		AA424029	Hs.81897	ESTs; Moderately similar to dynamin; int ESTs	5.4
50	115821	AA424038 AA427528		ESTs; Weakly similar to ZINC FINGER PROT	13
	115955		Hs.44198	Homo saplens BAC clone RG054D04 from 7q3	10
	116024	AA451748	Hs.83883	Human DNA sequence from clone 718J7 on c	6.8
~~	116108		Hs.28777	ESTs	6
55	116117		Hs.31575	SEC63; endoptasmic reticulum translocon ESTs	7.3 5.5
	116146	AA460701 AA489033	Hs.15423 Hs.62601	Homo sapiens mRNA; cDNA DKFZp586K1318 (	
		AA521472	Hs.71252	ESTs	5.5
		AA599463	Hs.306051		5.9
60	116401		Hs.59698	ESTs	7.9
		AA609219	Hs.39982	ESTs	9.2
	116587		Hs.121429		5.2
	116601 116684		Hs.45140 Hs.66095	ESTs ESTs	7.2
65	116722		0.00030	HSFIH32 Stratagene cat#937212 (1992) Hom	5.5
05	116766		Hs.95097	ESTs	5.8
	117453	N29568	Hs.108319	thyroid hormone receptor-associated prot	6.9
	117557		Hs.44532	diubiquitin	4.8
	117708	N45114	Hs.126280	ESIS	6.

		N52151	Hs.47447	ESTs	11.4
		N62339		heat shock 90kD protein 1; alpha	6.2
		N69207	Hs.203697		5.8
5		N70358 N89881	Hs.125180 Hs.44577	growth hormone receptor ESTs	7.1 6
,		N94303	Hs.55028	ESTS	9.3
	119107		Hs.63841	ESTs	6
	119126	R45175	Hs.117183	ESTs	17.9
	119271	T16387	Hs.65328	ESTs	6
10	119367	T78324	Hs.250895		5
	119721	W69440	Hs.48376	ESTs	15.4
	119741	W70205	Hs.43670	kinesin family member 3A	10.1
	119780	W72967 Z41078	Hs.191381 Hs.66035	ESTs; Weakly similar to hypothetical pro ESTs	5.3 4.8
15		AA173939		ESTs; Weakly similar to inner centromere	8.8
15	120294	AA190888		ESTs; Highly similar to NY-REN-62 antige	4.9
		AA236010	Hs.26613	Homo sapiens mRNA; cDNA DKFZp588F1323 (	14.7
		AA253400		turnor protein 63 kDa with strong homolog	5.6
••		AA261852	Hs.192905		4.9
20		AA280738	Hs.34892	ESTs	8.8
		AA282074	Hs.237323	ESTS	6.2 9.9
		AA292655 AA398246	Hs.96557 Hs.97594	ESTs	16.4
	121429		Hs.41167	ESTS	6.9
25	121503		Hs.290347		7.6
	121512	AA412105	Hs.193736		5.8
	121816	AA424814	Hs.48827	ESTs	4.6
		AA431302	Hs.98721	EST; Weakly similar to N-copine (H.saple	5.6
20		AA437311	Hs.98927	ESTs	5.7
30		AA446859 AA460158	Hs.99083	ESTs KIAA1028 protein	6.5 12.4
		AA460225	Hs.129030 Hs.99519	ESTs	5.1
		AA478539	Hs.104336		4.9
		AA485724	Hs.27413	ESTs	5.4
35		AA485957		Homo sapiens clone 25032 mRNA sequence	5
		AA495981	Hs.250830		4.7
		AA496252	Hs.105069		7.4
		AA609006 AA609200	Hs.111240	ESTs	9.1 4.7
40		AA609310	Hs.183691		4.8
10		AA609651	Hs.112742		7
	123938			damage-specific DNA binding protein 1 (1	5
	124178	H45996	Hs.97101	putative G protein-coupled receptor	6.8
. ~	124352	N21626	Hs.102406		10.2
45	124357	N22401	Hs.109370	yw37g07.s1 Morton Fetal Cochlea Homo sap ESTs	10.6 14.2
		N58172 R88992	Hs.174195		4.8
		W38419	110.174100	ESTS	4.7
	125992	W01626		za36e07.r1 Soares fetal liver spleen 1NF	5.1
50	126802	AA947601	Hs.97056	ESTs	5.1
	126812	Z36290	Hs.173933		4.6
		AA662913	Hs.190173		5
		AA507628	Hs.334390 Hs.70337		4.8 - 4.7
55		AI024352 AI457411	Hs.106728	Immunoglobulin superfamily; member 4 ESTs	4.7
55		AA828760	Hs.292059		4.8
		Al400362	Hs.265130		5
		AI039722	Hs.279009	ESTs	5.8
		AI088155	Hs.41296	ESTs; Weakly similar to unknown [H.saple	17
60		AA176446		ESTs; Weakly similar to hypothetical 43.	4.8
	128610		Hs.10247	activated leucocyte cell adhesion molecu ESTs; Weakly similar to KIAA0437 [H.sapl	7.9 8.1
	128625 128651		Hs.102032		6.5
	129068	AA215971	Hs.194431		5.2
65	129136		Hs.250723		5.1
	129171		Hs.7753	calumenin	5.8
	129229	AA211941		polyadenylate binding protein-interactin	5.8
		N27524		Cdo42 effector protein 3 ESTs	5.2
	129467	AA410311	Hs.44208	2919	5.1

	129564		Hs.75295	guanylate cyclase 1; soluble; alpha 3	16.3	
	129699 129821	AA459578 F11019	Hs.12017 Hs.12696	KIAA0439 protein; homolog of yeast ubiqu cortactin SH3 domain-binding protein	9.2 8.6	
	129823	X00948	Hs.105314	relaxin 2 (H2)	9.1	
5	129847	W46767	Hs.296178		5.4	
	129912	AA047344		ESTs; Highly similar to NY-REN-6 antigen	6.5	
	129958		Hs.1378	annexin A3	5.1	
	129977		Hs.1395	early growth response 2 (Krox-20 (Drosop	8.6	
10	130061			arginase; type It	7.4	
10	130241 130466	U78313 N21679	Hs.153203 Hs.180059	MyoD lamily inhibitor	4.9 5.8	
	130541	X05608		neurofilament; light polypeptide (68kD)	6.7	
	130619	AA477739	Hs.12532	ESTs	6.4	
	130925	N71935		multiple PDZ domain protein	7.9	
15	130938	AA013250	Hs.21398	ESTs; Moderately similar to PUTATIVE GLU	6.2	
	130971		Hs.301444	signal sequence receptor; gamma (translo	6.4	
	131066		Hs.22588	ESTs	5	
	131126			myotubularin related protein 2	6.4	
20	131310	J02960 AA253220	Hs.2551 Hs.27373	adrenergic; beta-2-; receptor; surface Homo sapiens mRNA; cDNA DKFZp564O1763 (	7.9 fc 0	
20	131487 131561	X59841		pre-B-cell leukemia transcription factor	7.6	
	131562	U90551	Hs.28777	H2A histone family; member L	5.1	
	131579		Hs.29088	ESTs	11	
	131629		Hs.238809		4.9	
25	131682	AA428368	Hs.30654	ESTs	4.8	
	131699		Hs.90421	ESTs; Moderately similar to IIII ALU SUB	6.5	
	131795		Hs.32317	Sox-like transcriptional factor	5.6	
	132053 132122		Hs.38085 Hs.40403	ESTs; Weakly similar to putative glycine Cbp/p300-interacting transactivator; wit	7.2 5.6	
30	132191	AA449431	Hs.288361		8	
50	132256	AA608856	Hs.431	murine leukemia viral (bmi-1) oncogene h	5.5	
		AA429478		ESTs; Highly similar to CGI-49 protein [	6.6	
	132533	AA021608	Hs.172510	ESTs	5.8	
	132572			signal recognition particle 72kD	6.2	
35	132581		Hs.52256	ESTs; Weakly similar to beta-TrCP protei	16	
	132700		Hs.5521 Hs.55220	ESTs	6.8 5.3	
	132701	AA279359 L41887		BCL2-associated athanogene 2 splicing factor; arglnlne/serlne-rich 7	7.8	
	132783			DEAD/H (Asp-Glu-Ala-Asp/His) box polypep	5.9	
40		X75535		peroxisomal famesylated protein	8	
	132939		Hs.61152	exostoses (multiple)-like 2	5.2	
	133142		Hs.65874	ESTs	5.2	
		U29589	Hs.7138	cholinergic receptor; muscarinic 3	10.3	
45		AA278852	Hs.30212	ESTS	5.8 4.9	
43	133520	M68941 X74331	Hs.73826 Hs.74519	protein tyrosine phosphatase; non-recept primase; polypeptide 2A (58kD)	13.1	
	133544		Hs.74624	protein tyrosine phosphatase; receptor t	4.6	
	133608		Hs.75207	glyoxalase I	4.8	
	133626	H75939	Hs.75277	Homo saplens mRNA; cDNA DKFZp586M141 (fi		
50	133633	D21262	Hs.75337	nucleolar phosphoprotein p130	6.3	
	133797	S66431	Hs.76272	retinoblastoma-binding protein 2	6	
	133928	N34096	Hs.7766	ubiquitin-conjugating enzyme E2E 1 (homo	5.4 5.2 -	
	134095	U47414 N89827	Hs.79069 Hs.80667	cyclin G2 RALBP1 associated Eps domain containing	6.5	
55	134321		Hs.8172	ESTs	7	
-	134453	X70683	Hs.83484	SRY (sex determining region Y)-box 4	4.7	
	134542	X57025	Hs.85112	insulin-like growth factor 1 (sornatomedi	7.7	
	134570	U66615	Hs.172280		6.4	
	134592	U82613	Hs.289104		5.4	
60	134654	W23625	Hs.8739	ESTs; Weakly similar to ORF YGR200c [S.c	5 5.4	
	134668	AA482319 Z49099	Hs.8752 Hs.89718	putative type II membrane protein spermine synthase	6.7	
	134806		Hs.169358		9.8	
	135066		Hs.93913	Interleukin 6 (Interferon; beta 2)	5.7	
65	135155	AA358268		ESTs; Moderately similar to transcriptio	4.9	
	135411	L10333	Hs.99947	reticulon 1	5.3	
	300023	M10098		AFFX control: 18S ribosomal RNA	4.6	
	300254	AW079607	Hs.55610	ESTs; Weakly similar to ZnT-3 [H.sapiens	7.8	
	300273	AW013907	ms.167531	ESTs; Moderately similar to predicted us	11.5	

		AW 13/040	115.155500	ED 13, Weakly Stillia to Illicolobolic-aco	0.0
	300566	H86709	Hs.326392	son of seveniess (Drosophila) homolog 1	5.8
	300578	Al989417	Hs.134289	ESTs	4.4
		AI239706		ESTs	7.9
5		AA039352		ESTs; Weakly similar to ORF YDL040c [S.c	4.5
				EGT - M. M. Jania to Oth Tocoroo jo.	5.2
		AW468066	Hs.24817	ESTs; Weakly similar to KIAA0986 protein	
		AI497778	Hs.20509	ESTs	6.4
	300810	AI076890	Hs.146847	ESTs	5.8
	300813	AA406411	Hs.208341	ESTs; Weakly similar to KIAA0989 protein	10.
10	300833	Al863068	He 105923	ESTs; Weakly similar to putative zinc fi	5.6
		AF109300	Hs.147924		6.7
			Hs.1852	ESTs	7.6
		AW136372			
		AA593373	Hs.293744		5.5
		AA947682	Hs.20252	ESTs; Weakly similar to Chain A; Cdo42hs	7
15	301042	Al659131	Hs.197733	ESTs	24.
	301242	AW161535	Hs.23782	ESTs	11.
	301254	Al049624	Hs.283390	EST cluster (not in UniGene) with exon h	4.3
		H29500	Hs.7130	ESTs; Moderately similar to N-copine [H.	4.3
		AA156879		ESTs; Weekly similar to ZINC FINGER PROT	6.6
20					5.7
20		Al802946	MS.442UG	ESTs; Weakly similar to match to ESTs AA	
		AW008475		EST cluster (not in UniGene) with exon h	6.8
	301689	Z44810		ESTs; Weakly similar to similar to C.ele	6.3
	301783	AL046347	Hs.83937	Homo sapiens PAC clone DJ1159O04 from 7p	6.2
		AI800004		ESTs; Weakly similar to MesP1 [M.musculu	8.5
25		R20002	Hs.8823	ESTs; Weakly similar to intrinsic factor	4.6
23		AF131855		Homo sapiens clone 25056 mRNA sequence	8.3
	201031	AFTOTOCO			38
		Al869666	Hs.123119		9.5
	302056	AI457532		ESTs; Moderately similar to ROSA26AS [M.	
		H05698		ESTs; Weakly similar to protein-tyrosine	5.8
30	302099	AL021397		ribosomal protein L34 pseudogene 1	8.8
	302147	AB022660		KIAA0437 protein	5.9
	302214	AJ001454	Hs.159425	Homo sapiens mRNA for testican-3	4.8
		Al128606	Hs.6557	zinc finger protein 161	4.3
	302358	D81150	Hs 322848	EST cluster (not in UniGene) with exon h	5.5
35		NM_004917	He 218368	EST cluster (not in UniGene) with exon h	26
50		AC003682		multiple UniGene matches	8.2
	000500	NM_000522		EST cluster (not in UniGene) with exon h	6.4
	302302	14M_000522		EST cluster (not in UniGene) with exon h	5
		AA425562	Hs.11065		4.8
40		AA343696	Hs.46821	ESTs; Weakly similar to putative [H.sapi	
40		AA508353		relaxin 1 (H1)	78
		N58545		histone deacetylase 3	8.5
	302970	AW118352		EST cluster (not in UniGene) with exon h	7.4
	302977	AW263124	Hs.315111	EST cluster (not in UniGene) with exon h	5.5
		AF199613		EST cluster (not in UniGene) with exon h	4.6
45		AF181352	Hs.111782	EST cluster (not in UniGene) with exon h	5.8
		AI571580	Hs.170307		4.3
		AA215297	Hs.61441	EST cluster (not in UniGene) with exon h	6.4
					6.6
		AL134164	Hs.145418		19
60		AA255977		ESTs; Highly similar to ubiquitin-conjug	
50		AA298471	Hs.326567	EST cluster (not in UniGens) with exon h	6.6
		AA758552	Hs.309497		6.8
	303525	AW518519	Hs.273294	ESTs	4.8
	303526	AA348111	Hs.96900	ESTs	12
		AA355607	He 309490	ESTs; Weakly similar to MMSET type I [H.	8.2
55		AW338520	Hs.242540		8.4
55		AW500106		EST cluster (not in UniGene) with exon h	4.9
				EST cluster (not in UniGene) with exon h	15
		D30891			6.3
	303702	AW500748		ESTs; Weakly similar to 73 kDA subunit o	
	303718	Al741397	Hs.114658		4.6
60		AA521510	Hs.145010		12
	303732	AW502405		ESTs; Weakly similar to tumor suppressor	4.3
	303735	AA707750	Hs.169055	ESTs; Wealdy similar to cis-Golgi matrix	5.4
		At017286	Hs.5957	EST cluster (not in UniGene) with exon h	5.3
	303753	AW503733	Hs.9414	ESTs	13
65		AI275850		EST cluster (not in UniGene) with exon h	7.8
	304053	R00493		translocase of inner mitochondrial membr	4.8
		N66373	Hs.27973		6
		AA668128	Hs.45207		5.
			Hs.251354	COTA	5.
	300/10	AI024916	ris.201304	Edia	J.,

300319 AW157646 Hs.153506 ESTs; Weakly similar to microtubule-acti

		Al364186		EST singleton (not in UniGene) with exon	7.3
		Al368665	Hs.31476	EST singleton (not in UniGene) with exon	5.4
		AI460004	Hs.31608	EST singleton (not in UniGene) with exon	8.1
_		Al613519		EST singleton (not in UniGene) with exon	5.5
5		Al863051	Hs.279815		4.4
	309116	AI927149	Hs.29797	ribosomal protein L10	4.5 7.4
		AW075342 AW205604	Hs.9271	EST singleton (not in UniGene) with exon ESTs; Weakly similar to !!!! ALU SUBFAMI	5
		AVI200004 AV921750	Hs.144871		5
10		Al685841	Hs.161354		11.8
10		AJ478629	Hs.158465		6.8
			Hs.145569		9.7
		A1734009	Hs.127699	EST cluster (not in UniGene)	104
		Al812775	Hs.145710		4.6
15		Al420227	Hs.149358	ESTs	72.9
		AW292180	Hs.156142	ESTs	7.6
		Al338013	Hs.140546 Hs.175162		9.2 4.5
	310639	AW269082 AW262580	Hs.147674		4.9
20		AVV202080 AI973051	Hs.224965		7.6
20			Hs.197698		41.
				ESTs; Weakly similar to Y38A8.1 gene pro	4.5
				ESTs; Moderately similar to !!!! ALU SUB	4.6
			Hs.23862	ESTs	5.9
25		AI824863	Hs.211420		4.8
		AI828254	Hs.271019		5.8
		Al682088	Hs.79375	ESTs	26.
		Al809519 .	Hs.27133	ESTs	6.4 7.4
30		AW025661	Hs.240090		7.4 4.6
30		AI682478 AA765470	Hs.13528 Hs.85092	ESTS	6.7
		AW014013	Hs.107056		5.3
		R16890	Hs.137135		5.6
			Hs.257482		4.3
35		AA759250		cytochrome b-561	11
	312182	AA834800	Hs.326263	EST cluster (not in UniGene)	18.
	312242	Al380207	Hs.125278		4.7
		C01387	Hs.127128		5.3
40		R46180	Hs.153485		6.2 4.8
40		AA847398	Hs.291997 Hs.293892		5.2
		R49353 R68651	Hs.144997		9.5
		C17785	Hs.182738	ESTe	6.3
		AA033609	Hs.239884		11.
45		Al695522	Hs.191271		4.7
		AI004377	Hs.200360		7
	312546	AI623511	Hs.118567		5.1
		AA976064	Hs.180842		8.5
		AA694607		EST cluster (not in UniGene)	10.
50		AA772279	Hs.126914		5 5.8
		Al813654	Hs.5957 Hs.278626	ESTs	7.7
		AA939266 H92571	Hs.234478		6.5
		AA836271	Hs.125830		4.8
55		A1079278	Hs.269899	ESTS	5.1
55		AA249018		EST cluster (not in UniGene)	7
		N36417	Hs.144928		6.3
	313188	AI801098	Hs.151500	ESTs	4.3
		AI039702	Hs.179573	collagen; type I; alpha 2	4.8
60		AA827805	Hs.124296		5 5.9
		Al200281	Hs.123910	ESIS	
	313325	Al420611 Al088120	Hs.127832		4.6 7.4
		AA745689	Hs.122329	ESTs; Weakly similar to similar to zinc	6.3
65		AA743000 AI261390	Hs.146085	FSTe	5.6
00		Al797301	Hs.5740	ESTS	5.9
		AW467376	Hs.129640	ESTs	4.3
	313569	Al273419	Hs.135146	ESTs; Weakly similar to ZK1058.5 [C.eleg	4.6
	313603	AW468119	Hs.267631	EST cluster (not in UniGene)	6.8

		AW295194		DKFZP434N126 protein	5.2 7.8
		AW468402 AA688292	Hs.254020 Hs.337786		4.4
		AA507227	Hs.6390	ESTS	8.1
5		AI753075	Hs.104627	ESTs	6.7
	313670	C16690		EST cluster (not in UniGene)	4.4
		W49823	Hs,104613		4.4
		AA861697	Hs.120591	EST cluster (not in UniGene)	13.4
10		Al161293 AA768553	Hs.74170	ESTs; Weakly similar to KIAA0525 protein	5.2
10		AMPORAGO	Me SECOR	ECTo	5.4
		AJ535895	Hs.221024	FSTs	4.3
		AI732100	Hs.187619	ESTs	13.6
	314123	AW245993	Hs.223394	ESTs	8.4
15		Al821895	Hs.193481		29.4
		AL138431	Hs.164243		4.8 5.7
		AL036001 AA743396	Hs.48376 Hs.189023		4.9
		AA732359	Hs.96264		4.4
20	314284	AA731431		EST cluster (not in UniGene)	6.4
	314305	Al280112	Hs.125232	ESTs	5.3
	314343	Al754701	Hs.328476	ESTs ESTs; Weakly similar to alternatively sp ESTs	6.2
	314530	AI052358	Hs.193726	ESTs	4.5
25		AW207206			17 8.9
25		AW502698 AI538226	Hs.118152 Hs.32976		9.4
		AA481027	Hs 109045	ESTs; Weakly similar to ORF YGR245c [S.c	8
	314864	AA493811	Hs.294068	ESTs	8
	314907	AA493811 A872225	Hs.222886	ESTs	19.3
30		AA548906			4.5
		AA521381			5.3
		AA524953 AA533447	Hs.293334	EST cluster (not in UniGene)	4.6 5.1
		AW292425	Hs.163484		15.5
35		AA876910	Hs.134427		20
	315073	AW452948		ESTs	5.3
	315084	Al821085		ESTs	8.2
	315214	AI915927 AI420753 AI985544 AI222165	Hs.34771	ESTS	5.4 5.1
40	315220	AM2U/53	HS.86731	ESIS ECTe	5.8
70	315280	A1222165	Hs 144923	FRTs	4.5
	315368	AW291563	Hs.104696	ESTS	8
	315369	AA764918	Hs.256531	ESTs	4.8
	315378	At263393	Hs.145008	ESTs	6.2
45	315379	Al378329	Hs.126629	ESIS	5.4 5.1
		AW293424 AA977935	Hs.75354 Hs.127274	ESTs FOTe	6.6
		AW003416	Hs.160604	ESTS	5.5
		R37257	Hs.184780	ESTs	8.1
50	315593	AW198103	Hs.158154	ESTs	9.9
	315634		Hs.220585	ESTs	7.8
		AW449285	Hs.313636	ESTS	8.9
		AI418055 AA744015	Hs.161160	EST cluster (not in UniGene)	5.1 6.1
55		T05558	Hs.156880	EST cluster (not in UniGene)	6.8
55		Al391470	Hs.158618		5.3
		AA744875	Hs.189413	ESTs	5
	315843	AA679430	Hs.191897	ESTs	5.7
60		AI800041	Hs.190555		9.2
60		AA764950 AA708016	Hs.119898 Hs.190389	FSTe	4.3 5.9
		AA693880	Hs.6947	EST cluster (not in UniGene)	6.7
		AW517542		ESTs	5.5
			Hs.213003	ESTs	5.1
65	316169	Al127483	Hs.120451	ESTs	8.2
		AA760894	Hs.153023	EST	17.1
	316491	AA766025 AW135854	Hs.186854 Hs.132458	EG1 FSTe	4.8
		AW015940	Hs.232234	ESTs	7.6
	J.0001				

		AA031215		ESTS, Weakly Similar to predicted distrig	5.1
	316905	AW138241	Hs.210846	ESTs	6.4
	317008	AW051597	Hs.143707	ESTs	4.4
		AA864968	Hs.127699	FSTe	11
5		AW445167	Hs,126036		13.
5		Promo IO	HS,120000	EGTS	8.7
	317224	D26760	Hs.93029	ESIS	
			Hs. 126594		8.7
		AA931245	Hs.137097	ESTs	11.
	317548	AI654187	Hs.195704	ESTs	14.
10	317651	AW292779	Hs.169799	ESTs	5.8
		AI733277	Hs.128321		5.4
				EST duster (not in UniGene)	11
	317850	N29974 AW295184 AI828602	MS, 102902	EST CUSSE (NOT IN CHICAGO)	
	317869	AW295184	HS.129142	ESTs; Weakly similar to DEOXYRIBONUCLEAS	
					5.3
15	317916	AI565071	Hs.159983	ESTs	7.7
	318239	AI085198	Hs.164226	ESTs	13.
		Al817736	Hs.182490	FSTs	8.2
			Hs.200942		4.6
					6
		R45530	Hs.1440	gamma-aminobutyric acid (GABA) A recepto	
20			Hs.194591		12.
	318464	Al151010	Hs.157774		4.3
	318524	AW291511	Hs.159066	ESTs	25.
	318540		Hs.274803	EST duster (not in UniGene)	7
	318501	AW206806	Hs.115325		4.8
25	21001		Hs.10177		5.5
43					5.7
	318646	AW175665	NS.278090		
	318657	Al493742	Hs.165210	ESIS	11
		W26276	Hs.136075		5.9
	318753	AA578265	Hs.7130	copine IV	5.5
30	319080	Z45131	Hs.23023	ESTs	16.
				EST cluster (not in UniGene)	4.6
					6.6
		R21054	Hs.180532		4.9
		D78808	Hs.283683	ECTA	8.2
35	210750	AA621605	Hs.117956		9.3
33	010700	AA40077E		ESTS	14.
			Hs.6295		12.
	319824				
		AA337642	HS.95262		5.1
					4.3
40	319964	T80579	Hs.290270	ESTs	5.8
	320076	Al659733	Hs.271593	ESTs	8.5
	320102	AW296219	Hs.115325	RAB7; member RAS oncogene family-like 1	9.8
		T99949	Hs.303428	EST cluster (not in UniGene)	9.8
					7.9
45		AF071202	Un 190996	ATP-binding cassette; sub-family C (CFTR	56.
43		R49889	D-04444	EST cluster (not in UniGene)	8.3
					5.4
	320464	Al089817	Hs.237146		
			Hs.159330	EST cluster (not in UniGene)	7
	320574	AL049443	Hs.161283	Homo sapiens mRNA; cDNA DKFZp586N2020 (	f 4.4
50	320576	ALC49977	Hs.162209	Homo sepiens mRNA; cDNA DKFZp564C122 (fr	6.7
		AW263086	Hs.118112	PSTs	6
		AF038966		secretory carrier membrane protein 1	13.
			Hs.71721	ESTs	6.2
		AI681006	115.71721	E015	9.3
		AW360847	Hs.16578	E513	
55			Hs.135904	ESIS	8.1
		D59945	Hs.65366	EST cluster (not in UniGene)	6
		AA633772	Hs.116796		9.2
	320918	AW195012	Hs.293970	ESTs	5
		H19732	Hs.247917	ESTs	5.9
60		AA018386	Hs.64341	ESTs	4.6
00		H52462		EST cluster (not in UniGene)	5.8
	321190	102402	115,103072	EST cluster (not in UniCens)	8.4
	321318	AB033041	MS.13/50/	EST cluster (not in UniGene)	
		AW372449		EST cluster (not in UniGene)	7.3
		AW297633	Hs,118498	ESTs	14.
65		H80483	Hs.46903	EST cluster (not in UniGene)	9.2
	321609	H86021	Hs.182538	ESTs; Weakly similar to hMmTRA1b [H.sapl	4.8
	321636	Al791838	Hs, 193465	ESTs	5.5
	321639	Al356352	Hs.108932	ESTs	4.6
	321644		Ha.237396		6.6
	W. 1044		. 1420. 550		-~

316854 AA831215 Hs.159066 ESTs; Weakly similar to predicted using

		AA233821	Hs.190173	EST cluster (not in UniGene)	4.6
		X91221	Hs.144465	EST cluster (not in UniGene)	5
		U29112	Hs.196151	EST cluster (not in UniGene)	6.2 4.6
5	3218//	N55158	Hs.189222 Hs.29468	EST cluster (not in UniGene)	4.8
5		AA746374	Hs.145010		8.2
	322007	AW410646			5.1
	322055	AL137646	Hs.146001	EST cluster (not in UniGene)	4.3
		AF085833		EST cluster (not in UniGene)	4.3
10	322221	AI890619	Hs.179662	nucleosome assembly protein 1-like 1	4.4
	322278	AF086283	11- 457004	EST cluster (not in UniGene)	5.8 22
	322303	MU/409	HS.15/001	EST custer (not in Uniciana)	4.4
	322407	AF143235	He 279819	EST cluster (not in UniGene) ESTs; Weakly similar to rabaptin-4 [H.sa EST cluster (not in UniGene)	7.2
15	322782	AA056060	Hs.202577	EST cluster (not in UniGene)	18.
	322811	AA782292	Hs.105872	EST cluster (not in UniGene) ESTs	6.9
	322818	AW043782	Hs.293616	ESTs	10.
		AI807883	Hs.180059		5
20		AI986306 AA081924	Hs.86149 Hs.124918	ESTs; Weakly similar to KIAA0969 protein	11. 7.1
20	322889	AA669253	Hs.124918 Hs.136076		4.5
	322982	AI351191	Hs.126430		6.6
	322994	AA422116	Hs.191461	ESTs	4.7
	323040	AA336609	Hs.10862	ESTs	6.9
25	323041	AL118747	Hs.26891	EST cluster (not in UniGene)	8.3
	323045	AA148950	Hs.188836	ESTs	4.6
	323048	AL118923	Hs.1/5110	EST cluster (not in Uniciene)	7.5 7.5
	323070	AA157720 AA157007	Hs.204330	EOTs ECTs	4.7
30	323097	Z44354	Hs.296261	guanine nucleotide binding protein (G pr	4.9
	323131	AA176982	Hs.270124	EST cluster (not in UniGene)	6.1
	323136	AL120351	Hs.30177	ESTs EST autor (not in UniGene) EST sustair (not in UniGene) EST sustair (not in UniGene) EST sustair (not in UniGene) EST autorities (not in UniGene) EST autorities (not in UniGene) EST cluster (not in UniGene) EST autorities (not in UniGene) EST autorities (not in UniGene) EST autorities (not in UniGene)	4.3
	323175	AI827137	Hs.336454	ESTs	6.2
35	323218	AF131846	Hs.13396	Homo sapiens clone 25028 mRNA sequence	
33			Hs.21906 Hs.293960		10.
	323230	AI829770	Hs.190642	FSTe	7.6
					7.6
	323287	AA639902	Hs.104215	ES1 a EST a	24.
40	323335	AI655499	Hs.161712	ESTs	14.
	323341	AL134875	Hs.108646	ESTs	5.3
	323362	AL135067	Hs.117182	ESTS Moderately similar to IDVDI IVATE DE	6.1 8.5
	323466	AIR26801	Hs 300700	ESTs, Moderately Similar to [FTHOVATE DE	4.5
45	323507	H71721	Hs.128387	ESTs	4.4
	323545	AI814405	Hs.224569	ESTs	5.8
	323623	AA314280	Hs.146589	EST cluster (not in UniGene)	5
	323663	AW263526	Hs.243023	ESTs	7.7
50	323691	AA317561	Hs.145599		5.9 8.2
50		AA740405 AA337621	Hs.108806 Hs.137635		6
		AA354940	Hs.145958		10.
		Al636775		ESTs	5.4
		AA367032	Hs.217882	ESTs	5.8
55	323997	AA844907	Hs.274454	EST cluster (not in UniGene)	4.4
		AW177009		EST cluster (not in UniGene)	4.6 11
	924305	AL046575 Al146686	Hs.143691	ESTs	13.
			Hs.192524		6.8
60	324307	AA627642	Hs.4994	transducer of ERBB2; 2 (TOB2)	4.9
	324330	AA884766		EST cluster (not in UniGene)	4.3
	324385	F28212	Hs.284247	EST cluster (not in UniGene) EST cluster (not in UniGene)	4.7
	324430	AA464018	Hs.184598	EST Cluster (not in UniGene)	13. 7.6
65	324452	AW014022 AW501974	Ms.170953	ECTo	5.6
33	324697	AW016378	Hs 292934	ESTs	24.
	324617	AA508552	Hs.195839	ESTs	54
	324618	AI346282	Hs.87159	ESTS	4.6
	324620	AA448021	Hs.94109	EST duster (not in UniGene)	5.7

	324626	Al685464		ESTs	9	
		Al694767	Hs.129179	ESTs	22	
	324876		Hs.112451		4.9	
	324891			ESTs; Weakly similar to Pro-a2(XI) [H.sa	10.6	
5			Hs.257339		10.2	
			Hs.163440		5.5	
		A1739168	Hs.116467	EST cluster (not in UniGene)	7.2 34.4	
		Al557019	Hs.292437		4.8	
10		AA578904 Al279919		ESTS; Moderately similar to !!!! ALU SUB	7.9	
10		AA612626	Hs 144871	EST cluster (not in UniGene)	5.2	
		Al334367	Hs.159337	ESTs	7.6	
		Al619924	Hs.14553		12.6	
		AI692552		ESTs	6.5	
15	324845	AA361016	Hs.337533		4.5	
		Al564134		KIAA0853 protein	4.4	
		AI741633	Hs.125350		6.5	
		AA613792	11- 00000	EST cluster (not in UniGene)	5.1 7.1	
20		AA401863	Hs.22380	CH.20_hs gij6552458	9.6	
20	326816 326997			CH.20_1is gij0002406 CH.21_hs gij5867660	4.6	
	327098			CH.21_hs gij6682516	4.3	
	328492			CH.07_hs gi[5868455	5.8	
	329362			CH.X_hs gi[5968837	4.3	
25	329929			CH.16_p2 gl[6165201	5.5	
	329960			CH.16_p2 gi[5091594	7.6	
	330020			CH.16_p2 g  6671887	6	
	333211			CH.05_p2 gi 6013592	12.6	
20	330384	M23263		androgen receptor (dihydrotestosterone r	Antigen, Prostate Specific, Alt. Splice	13.8
30	330430	HG2261-HT23	bii H- naana	Hs.321110 guanine nucleotide binding protein 4	R Antigent, Prostate opecino, Air. Spice	10.0
		U31362 U39840	H82293007	hepatocyte nuclear factor 3; alpha	4.9	
		AA319514	Hs.30732	FSTs	6	
		AA037415	Hs.20999	ESTs	5.5	
35		AA056557	Hs.6759	ESTs	5.1	
	330705	AA102571	Hs.157078	ESTs	11.7	
	330706	AA121140		ESTs; Moderately similar to kynurenine a	14.5	
	330712	AA167269	Hs.52620	ESTs	5	
40	330725	AA252033	Hs.24052	ESTs; Weakly similar to !!!! ALU SUBFAMI	7.2	
40		AA281092	Hs.35254	ESIS	4.9 18.5	
		AA449677		Human DNA sequence from clone 437M21 on FK506-binding protein 3 (25kD)	4.3	
		AA450200 AA479114	Hs.11356		5.8	
		D60374	113.11000	EST	4.6	
45	330892	AA149579	Hs.91202		15.3	
	330949	H01458	Hs.142896		10.3	
	330977	H20826	Hs.315181	ESTs	4.4	
		N24619	Hs.108920		11.8	
		R36671	Hs.14846		11.6	
50		R51361	Hs.268714	ESTs	4.8 13	
		R82331	Hs.268838 Hs.168439		4.9	
	331195	T64447 AA262999	Hs.300141		4.6 -	
	221221	AA278355	Hs.87929		6.1	
55	391337	AA287662	Hs.118630		9.2	
55		AA400596	Hs.88143		9.9	
		AA416979	Hs.81897		4.3	
		AA454543	Hs.43543		4.6	
		F10802		ESTs; Moderately similar to IIII ALU SUB	4.9	
60		H77381	Hs.41223		7.5	
		N21680	Hs.43455	ESTs	5.4 6.5	
		N27154	Hs.44076	ESTs	12.5	
		N32912 N34357	Hs.93817	ESTs; Weakly similar to hypothetical 43.	4.6	
65		N62780	Hs.48703		9.2	
03		N92352	Hs.5472	ESTs	4.6	
	331659	W48868	Hs.334305		8.7	
	331696	Z38907	Hs.65949	KIAA0888 protein	10.3	
	331811	AA404500	Hs.187958	ESTs	4.8	
				121		
				131		

	331848	AA417039	Hs.98268	signal recognition particle 72kD	7.5
	331873	AA429445	Hs.98640	ESTs	6.5
	331889		Hs.98802	Homo sapiens Chromosome 16 BAC clone CIT	33.6
~	331967	AA480158	Hs.99589	KIAA1028 protein	6.8
5	331974		Hs.105322		5.3 10.8
		AA490831 AA599477	Hs.201591 Hs.291156		4.4
	332173		Hs.100725	ESTs ESTs	5.5
	332247		113.100720	ESTs	14.2
10	332249		Hs.194140		7.2
	332325	T79428	Hs.339667	ESTs	5.6
	332396	AA340504		ESTs; Weakly similar to similar to human	21.2
	332434			transcription factor 4	15.3
		N95495	Hs.56729	ESTs; Highly similar to GTP-binding prot	7.1
15	332522			glutathione S-transferase theta 2	6.8
		AA281753 M31682	Hs.17731 Hs.19280	inositol 1;4;5-triphosphate receptor; ty inhibin; beta B (activin AB beta polypep	5.8 5.5
		M99487		folate hydrolase (prostate-specific memb	38.1
	332538		Hs.20991	ESTs	6.5
20	332546		Hs.22587	solute carrier family 35 (UDP-galactose	4.8
	332594	AA279313	Hs.32951	methyl CpG binding protein 2	5.6
	332610	AA412405	Hs.40513	ESTs; Weakly similar to BETA GALACTOSIDA	5.6
	332661	N95742	Hs.6390	ESTs	6.9
~ ~	332697		Hs.75725	carboxypeptidase E	24.3
25		D26070	Hs.79306	inosilol 1;4;5-triphosphate receptor, ty	9.9
	332718		Hs.79630	v-myc avian myelocytomatosis viral oncog	5.6 5
		R72029 AA233258	Hs.83428	synaptophysin-like protein ESTs; Weakly similar to D1007.5 (C.elega	4.5
	332797	MH233230		CH22_FGENES.6_2	30.8
30	332798			CH22_FGENES.6_5	66.8
	332799			CH22_FGENES.6_6	19.8
	332933			CH22_FGENES.38_7	5.6
	332980			CH22_FGENES.54_1	5.5
25	332984			CH22_FGENES.54_6	4.9
35	333168			CH22_FGENES.94_1 CH22_FGENES.94_2	4.7
	333169 333452			CH22_FGENES.157_1	4.8
	333456			CH22 FGENES.157_5	4.3
	333458			CH22 FGENES.157_7	4.6
40	333611			CH22_FGENES.217_6	4.7
	333621			CH22_FGENES.219_5	5.5
	333814			CH22_FGENES.282_2	7.1
	333849			CH22_FGENES.290_8 CH22_FGENES.303_5	6.2 4.3
45	333949 333951			CH22_FGENES.303_7	4.9
43	333955			CH22 FGENES.303_11	5.6
	334150			CH22 FGENES.339.1	5.1
	334223			CH22_FGENES.380_4	20.3
	334297			CH22_FGENES.372_3	9.4
50	334443			CH22_FGENES.387_2	4.8
	334444			CH22_FGENES.387_4	5.6 13.1
	334447			CH22_FGENES.387_7 CH22_FGENES.405_11	13.1 5.4 -
	334570 334749			CH22_FGENES.427_1	5.3
55	334777			CH22_FGENES.430_9	4.7
55	334960			CH22_FGENES.465_29	5.2
	335179			CH22_FGENES.504_9	8.8
	335293			CH22_FGENES.527_6	4.7
	335550			CH22_FGENES.578_11	5.1
60	335581			CH22_FGENES.581_19	5.7
	335586 335809			CH22_FGENES.581_25 CH22_FGENES.617_6	4.3 6.2
	335810			CH22_FGENES.617_7	5.8
	335822			CH22_FGENES.619_7	7.1
65	335824			CH22_FGENES.619_11	8.5
	335853			CH22_FGENES.626_5	4.3
	<b>3</b> 35886			CH22_FGENES.632_4	4.3
	336034			CH22_FGENES.678_5	6.8
	336441			CH22_FGENES.827_7	7.6

	336624	CH22_FGENES.6-3	43.3
	336625	CH22_FGENES.6-4	37.9
	336679	CH22_FGENES.43-7	5.3
	337577	CH22_C65E1.GENSCAN.8-1	4.9
5	338255	CH22_EM:AC005500.GENSCAN.276-3	13.4
	338260	CH22_EM:AC005500.GENSCAN.279-10	4.6
	338561	CH22_EM:AC005500.GENSCAN.421-5	4.6
	338562	CH22_EM:AC005500.GENSCAN.421-6	4.3
	338759	CH22_EM:AC005500,GENSCAN.517-6	5.1
10	338763	CH22_EM:AC005500,GENSCAN.517-16	5.5
	338764	CH22_EM:AC005500.GENSCAN.517-17	7.1

PCT/US01/32045 WO 02/30268

TABLE 3A shows the accession numbers for those primekeys lacking unigeneID's for Table 3. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers for sequences comprising each cluster are listed in the "Accession" column.

Unique Eos probeset identifier number

10

```
Pkey:
CAT number:
                              Gene cluster number
         Accession:
                              Genbank accession numbers
15
         Pkey CAT number
                                               Accession
         123619 371681 1
                                               AA602964 AA609200
                                              Z24878 AA494096 F13654 AA494040 AA143127
         116722 143512 1
         103677 41847 1
                                               Z83806 AJ132091 AJ132090
20
         125992 1589048_1
                                               H48372 W01626
         109342 genbank_AA213620
                                               AA213620
         125154 genbank_W38419
                                               W38419
         101447 entrez M21305
                                               M21305
         124357 genbank_N22401
                                               N22401
25
                                               AA136590
         108910 genbank_AA136590
322278 47271_1
                                            HISBORY INCUREOUS VERSION
AIRS (1056 AWRYSIAH AASSAGE AIRS (1881 AAB57439 AAB40756 AAB50339
AWIT7006 AI831610
AAB84766 AWRYSIA AAB52975 AA447312
AIB85845 AWRYSISS AAR5387 AAB25142
AF196815 AF108766
                                               W69304 AF086283 W69200
         315084 350969_1
         324019 262792 1
         324330 300543 1
30
         324626 336411_1
         303029 37699 1
         324804 396093 1
                                             Al692552 Al393343 Al800510 Al377711 F24263 AA661876
         324981 376239_1
                                              AA613792 AW182329 T05304 AW858385
         329362 c_x_hs
35
         336624 CH22_4071FG_6_3_
         336625 CH22 4072FG 6_4
         336679 CH22 4157FG 43_7
         338255 CH22_6856FG__LINK_EM:AC00
         338260 CH22_6863FG__LINK_EM:AC00
40
         329929 c16_p2
         329960 c16_p2
        339561 CH22_7294FG_LINK_EM:AC00
339562 CH22_7295FG_LINK_EM:AC00
338759 CH22_7591FG_LINK_EM:AC00
         338763 CH22 7585FG LINK EM:AC00
         338764 CH22 7586FG LINK EM:AC00
         333168 CH22_400FG_94_1_LINK_EM:A
         333169 CH22_401FG_94_2_LINK_EM:A
         333452 CH22 702FG 157 1 LINK EM:
50
        333456 CH22_706FG_157_5_LINK_EM:
         333458 CH22_708FG_157_7_LINK EM:
         333611 CH22 872FG 217 6 LINK EM:
         333621 CH22 882FG 219 5 LINK EM:
         333814 CH22 1083FG 282 2 LINK EM
55
        333849 CH22_1118FG_290_8_LINK_EM
         335179 CH22 2515FG 504 9 LINK EM
         333949 CH22_1225FG_303_5_LINK_EM
         333951 CH22 1227FG 303 7 LINK EM
         333955 CH22_1231FG_303_11_LINK_E
335293 CH22_2635FG_527_6_LINK_EM
60
         326816 c20 hs
         326997 c21 hs
         335550 CH22_2905FG_576_11 LINK E
         335581 CH22_2938FG_581_19_LINK_E
65
         335586 CH22 2944FG 581 25 LINK E
```

328492 c\_7\_hs 335809 CH22 3181FG 617 6 LINK EM 335810 CH22\_3182FG\_617\_7\_LINK\_EM 335822 CH22\_3195FG\_619\_7\_LINK\_EM 5 335824 CH22\_3197FG\_619\_11\_LINK\_E 335853 CH22\_3228FG\_626\_5\_LINK\_EM 335886 CH22 3261FG 632 4 LINK EM 330020 c16\_p2 330211 c\_5\_p2 10 337577 CH22 5864FG LINK C65E1.G 307848 Al364186 332797 CH22 13FG 6 2 LINK C4G1.G 332798 CH22 14FG 6 5 LINK C4G1.G 332799 CH22\_15FG\_6\_6\_LINK\_C4G1.G 15 334150 CH22\_1429FG\_339\_1 LINK\_EM 332933 CH22\_154FG\_38\_7\_LINK\_C20H 332980 CH22\_204FG\_54\_1\_LINK\_EM:A 332984 CH22\_208FG\_54\_6\_LINK\_EM:A 334223 CH22\_1507FG\_360\_4\_LINK\_EM 334297 CH22\_1588FG\_372\_3\_LINK\_EM 20 327098 c21\_hs 334443 CH22\_1742FG\_387\_2\_LINK\_EM 334444 CH22\_1743FG\_387\_4\_LINK\_EM 334447 CH22\_1746FG\_387\_7\_LINK\_EM 334570 CH22\_1675FG\_405\_11\_LINK\_E 334749 CH22\_2061FG\_427\_1\_LINK\_EM 25 334777 CH22 2089FG 430 9 LINK EM 335034 CH22 3419FG 678 5 LINK DJ 334960 CH22-2281FG 465 29 LINK E 30 336441 CH22\_3861FG\_827\_7\_LINK\_DJ U39840 NM\_004496 AW135907 BE087458 BE087567 AA177116 AW195705 AW750756 AI811008 AI694151 330551 9851 2 BE348594 AW971075 Al347950 Al201455 Al073898 AA652680 AA613671 Al318964 AA607550 AA693692 AI032599 AA991871 AI289801 AW948974 T74639 AA532907 AW949173 BE379594 Al192455 AL039862 Al744012 Al761735 AW243181 Al743687 Al928223 Al423022 Al627855 330786 53973 3 35 AI636059 AI651571 AWR02044 AI826995 AI431733 AI539125 AA863056 AW270910 AI768930 AW008835 AW615183 AW591147 Al695294 Al672106 AA506358 Al308060 AA011556 AA962437 Al935488 BE219625 AI004356 AW151394 AI218466 N66178 AI419784 AW242519 AW946907 D60374 AA989263 AI698799 AA470480 Al824167 AA669097 AA513815 AA026798 AA676526 AA704429 AA704269 AW118292 AA579216 N58172 332247 372969\_1 AW579842 BE156562 BE156690 BE156489 BE081033 AK001559 BE149402 M85387 AW367811 AW367798 40 332396 20265\_1 R17370 Al908047 AA382932 R58449 H18732 AA371231 AW962899 AA713530 AW892946 R53463 H11063 AW068542 Z40761 BE176212 BE176155 W23952 W92183 AW374883 AA303497 AW954769 AA036808 BE168063 AW382073 AW382085 AL041475 H80748 Al078161 BE463983 Al805213 Al761264 W94865 N94502 Al623772 Al419532 Al810302 Al634190 AW002516 AW150777 Al352312 Al367474 AW204807 45 Al675502 Al937026 AW134715 BE328451 Al123157 Al560020 Al900745 Al608631 Al248873 AA742484 AW051635 H18646 AI245045 AA507111 AI640510 AI925594 AA115747 AA143035 AA151106 AK001764 BE313896 AA380199 AA380151 AA194996 AW118089 AA495871 AW975219 AW085598 332781 32044 1 Al378909 AW992310 AW992409 Al911857 AA657643 Al804471 Al242589 Al623968 R09556 Al129100 Al206500 AA680094 AA677784 Al023178 Al277519 AA424742 Al240654 AA232846 Al804273 Al382376 50 AA001729 W90750 BE090656 AW295015 Al674596 Al431734 AM20517 AW769185 Al128355 Al192474 AI820001 AA001929 AA706925 AJ076676 AI499119 AI200493 AI695919 AI376217 W69195 W69261 AW305099 W90320 BE048357 Al658856 AA838534 AA233258 AI753393 AA709227 Al674387 AI872616

TABLE 3B shows the genomic positioning for those primekeys lacking unigene ID's and accession numbers in Table 3. For each predicted exon, we have listed the genomic sequence source used for prediction. Nucleotide locations of each predicted exon are also listed.

Play: Unique number corresponding to an Eos probeset
Ref: Sequence source. The 7 digit numbers in this column are Genbank Identifier (QI) numbers. "Ourhern L et al." refers to the
publication entitled "The DNA sequence of human chromosome 22." Dunham L et al., Nature (1999) 402-489-485.
Incluses both Astend form which cores were predicted.

5

O Strand: Nt_position:		ion: in	Indicates DNA strand from which exprs were predicted. Indicates nucleotide positions of predicted exons.				
5	Pkey	Ref	Strand	Nt_position			
,	333611	Dunham, I. et.al.	Plus	6548369-6548507			
	333621	Dunham, I. et.al.	Plus	8597414-8597560			
		Dunham, I. et.al.		7894165-7894252			
		Dunham, I. et.al.		8018323-8018472			
0		Dunham, I. et.al.		8589634-8589791			
		Dunham, I. et al.		8592501-8592637			
		Dunham, I. et.al.		8597414-8597560			
		Dunham, I. et.al.		10529221-10529854			
		Dunham, I. et al.		13420934-13421058			
5		Dunham, I. et.al.		14298981-14299058			
,		Dunham, I. et.al.		14306433-14306492			
		Dunham, I. et.al.		14308764-14308824			
		Dunham, I. et al.		14994868-14994943			
		Dunham, I. et.al.		16259596-16260166			
0		Dunham, I. et.al.		21634405-21634526			
•		Dunham, I. et.al.	Plus	24976198-24976334			
		Dunham, I. et al.		24990333-24990497			
		Dunham, I, et.al.	Plus	26310772-26310909			
		Dunham, I. et.al.	Plus	26314767-26314849			
5	335822	Dunham, I. et.al.	Plus	26364087-26364196			
,		Dunham, I. et.al.	Plus	26376860-26376942			
		Dunham, I. et.al.	Plus	26934235-26934364			
		Dunham, I. et.al.	Plus	29014404-29014590			
		Dunham, I. et.al.	Plus	34187606-34187663			
0		Dunham, I. et.al.	Plus	595377-595678			
J		Dunham, I. et.al.		15458919-15459257			
		Dunham, I. et.al.		216964-216798			
		Dunham, I. et.al.		232147-231974			
		Dunham, I. et.al.		232.447-231974 232.421-232307			
5		Dunham, I. et.al.		2035790-2035681			
,		Dunham, I. et.al.		5136165-5136019			
		Dunham, I. et.al.		2632606-2632457			
		Dunham, I. et.al.		3729896-3729788			
		Dunham, I. et.al.		3730864-3730767			
0		Dunham, I. et.al.		5136165-5136019			
U		Dunham, I. et.al.	Minus	2631933-2631797			
		Dunham, I. et.al.	Minus	5143942-514380 <del>6</del>			
		Dunham, I. et.al.		12734365-12734269			
		Dunham, I. et.al.	Minus	16090866-16090106			
5							
,		Dunham, I. et.al. Dunham, I. et.al.		20160968-20160795 22316408-22316275			
		Dunham, I. et.al. Dunham, I. et.al.		24689714-24668659 26614629-26614506			
0		Dunham, I. et.al. Dunham, I. et.al.		227714-227577			
U		Dunham, I. et.al.		229124-229024			
				2035790-2035681			
		Dunham, I. et.al.		15242294-15242231			
		Dunham, I. et.al.		22311966-22311856			
5		Dunham, I. et.al.		22312594-22312465			
J		Dunham, I. et.al.		26582475-26582199			
		Dunham, I. et.al.		26628148-26628009			
	338764	Dunham, I. et.al.	minus	26641232-26641101			

	329960	5091594	Minus	1031-1162
	329929	6165201	Minus	156410-156553
	330020	6671887	Plus	172397-172491
	326816	6552458	Plus	198354-198436
5	326997	5867660	Minus	71389-72147
	327098	6682516	Minus	1061684-1062361
	330211	6013592	Plus	59158-59215
	328492	5868455	Minus	46094-46241
	329362	5868837	Minus	65688-68173
10				

TABLE 4: shows a preferred subset of the Accession numbers for genes found in Table 3 which are differentially expressed in prostate tumor tissue compared to normal prostate tissue.

Pkay: Unique Eos probeset identifier number
ExAcon: Exemplar Accession number, Genbank accession number
Unique number
Unique number

Unigene Title: Unigere gene title
R1: Ratio of tumor to normal body tissue

	Pkey	ExAcen	UnigenelD	Unigene Title	R1
15					
		HG4020-HT42			10.5
		U75272	Hs.1867	progastricsin (pepsinogen C)	10.6
		X02544	Hs.572	arasamucaid 1	22.6
		AA236476	Hs.22791	ESTs; Weakly similar to transmembrane pr	10.3
20		AA282138	Hs.11325		14
		AA419461	Hs23317	ESTs	10.9
		AA156790	Hs.262036		15.3
		F01811		ESTs; Moderately similar to voltage-gate	10.8
05		T23855		KIAA1028 protein	10.8
25		Z38595	Hs.125019	ESTs; Highly similar to KIAA0886 protein	21.3
		AA460158		KIAA1028 protein	12.4
		N21626	Hs.102406		10.2
		Al659131	Hs.197733		24.9
30		Al869666	Hs.123119		36.8
30		NM_004917		EST cluster (not in UniGene) with exon h	26.8
		AA508353		relaxin 1 (H1)	78.8
		AA255977		ESTs; Highly similar to ubiquitin-conjug	19.5
		AW503733	Hs.9414	ESTS	13
35		Al420227 Al655662	Hs.149358 Hs.197698		72.9 41.3
53		Al682088	Hs.79375		26.4
		AA759250		cytochrome b-561	11
		AA033609	Hs.239884		11.2
		AA861697		EST duster (not in UniGene)	13.4
40		AI821895	Hs.193481		29.4
70		Al672225	Hs.222886		19.3
		AW292425	Hs.163484		15.5
		AA876910	Hs.134427		20
		Al654187	Hs.195704		14.2
45		AW295184		ESTs: Weakly similar to DEOXYRIBONUCLE	
		Al949409	Hs.194591		12.3
		AW291511	Hs.159066		25.9
	319080		Hs.23023		16.9
		AA460775	Hs.6295	ESTs	14.3
50		AF071202		ATP-binding cassette; sub-family C (CFTR	56.2
	321441	AW297633	Hs.118498		14.7
	322303	W07459	Hs.157601	EST cluster (not in UniGene)	22
		AA056060		EST cluster (not in UniGene)	18.4
	322818	AW043782	Hs.293616	ESTs	10.7
55	323287	AA639902	Hs.104215	ESTs	24.7
	324603	AW016378	Hs.292934	ESTs	24.2
	324617	AA508552	Hs.195839	ESTs	54
		Al694767	Hs.129179		22
	324691	Al217963		ESTs; Weakly similar to Pro-a2(Xi) [H.sa	10.6
60		AA641092	Hs.257339		10.2
		Al557019	Hs.116467		34.4
	330211			CH.05_p2 gi[6013592	126
				O Antigen, Prostate Specific, Alt. Splice	13.8
		AA121140		ESTs; Moderately similar to kynurenine a	14.5
65		AA449677		Human DNA sequence from clone 437M21 o	
		AA149579	Hs.91202		15.3
	330949	H01458	Hs.142896	ESIS	10.3

	331099	R36671	Hs.14846	ESTs	11.6
	331151	R82331	Hs.268838	ESTs	13
	331889	AA431407	Hs.98802	Homo sapiens Chromosome 16 BAC clone	CIT 33.6
	332247	N58172		ESTs	14.2
5	332396	AA340504		ESTs; Weakly similar to similar to human	21.2
	332533	M99487	Hs.325825	folate hydrolase (prostate-specific memb	38.1
	332697	T94885	Hs.75725	carboxypeptidase E	24.3
	332797			CH22_FGENES.6_2	30.8
	332798			CH22_FGENES.6 5	66.8
10	332799			CH22_FGENES.6_6	19.8
	334223			CH22_FGENES.360_4	20.3
	336624			CH22 FGENES.6-3	43.3
	338625			CH22 EGENES 6-4	37.9

TABLE 4A shows the accession numbers for those primekeys lacking unigeneID's for Table 4. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers for sequences comprising each cluster are listed in the "Accession" column.

5

30

10 Pkev: Unique Eos probeset identifier number CAT number: Gene cluster number Accession: Genbank accession numbers Pkey CAT number 15 Accession 336624 CH22 4071FG 6 3 336625 CH22\_4072FG\_6\_4\_ 330211 c\_5\_p2 20 332797 CH22\_13FG\_6\_2\_LINK\_C4G1.G 332798 CH22 14FG 6 5 LINK C4G1.G 332799 CH22\_15FG\_6\_6\_LINK\_C4G1.G 334223 CH22\_1507FG\_360\_4\_LINK\_EM 332247 372969\_1 AA669097 AA513815 AA626798 AA676526 AA704429 AA704269 AW118292 AA579216 N58172 25 332396 20265 1 AW579842 BE156562 BE156690 BE156489 BE081033 AKC01559 BE149402 M95387 AW367811

ANDTHAS DE EISANDE DE EISANDE DE EISANDE BEGEN DA BEGEN EISA AND DES DE LANGE, MISSON ANNOTATE IN ANKETTRE HITCH AND STANDE DE EISANDE MISSON ANKETTRE HITCH AND STANDE HITCH AND AND STANDE HITCH AND STANDE HITC

TABLE 4B shows the genomic positioning for those primekeys lacking unigene ID's and accession numbers in Table 4. For each predicted exon, we have listed the genomic sequence source used for prediction. Nucleotide locations of each predicted exon are also listed.

10	Pkey: Ref: Strand: Nt_position	Secuer DNA se Indica	ice source. To equence of hu tes DNA stra	nesponding to an Eos probesel n 7 (cf) transbors in the comma and deshank identifier (01) numbers. "Ourhant Let al." refers to the publication entitle not reference in 20 cmular et al., Nature (1999) 462-469-465, p position of prediction of the community of the	ed "T
15	Pkey	Ref	Strand	Nt_position	
	332797 332798	Dunham, I. et.al. Dunham, I. et.al.	Minus Minus	216964-216798 232147-231974	
	332799	Dunham, I. et.al.	Minus	232421-232307	
20	334223	Dunham, I. et.al.	Minus	12734365-12734269	
	338624	Dunham, I. et.al.	Minus	227714-227577	
	336625	Dunham, I. et.al.	Minus	229124-229024	
	330211	6013592	Plus	59158-59215	

## TABLE 5: 1170 GENES UP-REGULATED IN PROSTATE CANCER COMPARED TO NORMAL ADULT TISSUES

- 5 Table 5 shows 1170 genes up-regulated in prostate cancer compared to normal adult tissues. These were selected from 59680 probesets on the Affymetrix/Eos Hu03 GeneChip array such that the ratio of "average" prostate cancer to "average" normal adult tissues was greater than or equal to 3.44. The "average" prostate cancer level was set to the 85th percentile amongst 73 prostate cancers. The "average" normal adult tissue level was set to the 85th percentile
- 10 amongst 162 non-malignant tissues. In order to remove gene-specific background levels of non-specific hybridization, the 7.5<sup>th</sup> percentile value amongst the 162 non-malignant tissues was subtracted from both the numerator and the denominator before the ratio was evaluated.

15	Pkey: ExAcon: UnigenelD: Unigene Title: R1:		Unique Eos probeset identifier number Exemplar Accession number, Genbank accession number Uniquen enumber Uniquen egnes title				
			satio of tumor to				
			MANU OF IURING IN	Hoffina ussus			
20	Pkey	ExAcon	UnigenelD	Unigene Title .	R1		
	446057	Al420227	Hs.149358	ESTs. Weakly similar to A46010 X-linked	86.42		
	400302	N48056	Hs.1915	folate hydrolase (prostate-specific memb	66.46		
	414569	AF109298	Hs.118258	prostate cancer essociated protein 1	58.36		
25	417407	AA923278	Hs.290905	ESTs, Weakly similar to protease [H.sapi	56.16		
	431579	AW971082	Hs.222886	ESTs, Weekly similer to TRHY_HUMAN TRICH	53.38		
	409361	NM_005982	Hs.54416	sine oculis homeobox (Drosophila) homolo	48.28		
	409731	AA125985	Hs.56145	thymosin, beta, identified in neuroblast	45.24		
	400298	AA032279	Hs.61635	six transmembrane epithelial antigen of	43.48		
30	420154	AI093155	Hs.95420	JM27 protein	41.12		
50	433466	AA508353	Hs.105314	relaxin 1 (H1)	39.88		
	400296	AA305627	Hs.139336	ATP-binding cassette, sub-family C (CFTR	38,42		
	400292	AA250737	Hs.72472	ESTs	38,00		
	432887	Al926047	Hs.162859	ESTs	36.48		
35	439176	Al446444	Hs.190394	ESTs, Weakly similar to B28096 line-1 pr	36,45		
55	430722	AW968543	Hs.203270	ESTs, Weakly similar to ALU1_HUMAN ALU S	33.20		
	437052	AA861697	Hs.120591	ESTs	33.02		
	418398	AI765805	Hs.26691	ESTs	32.68		
	434036	A/659131	Hs.197733	hypothetical protein MGC2849	32.44		
40	407709	AA456135	Hs.23023	ESTs	32.10		
-10	426747	AA535210	Hs.171995	kallikrein 3, (prostate specific antigen	31.80		
	407168	R45175	110.171000	ESTs	31.72		
	440260	Al972867	Hs.7130	copine IV	30.52		
	421513	X00949	Hs.105314	relaxin 1 (H1)	30.10		
45	42 1513	N90470	Hs.203697	ESTs, Weakly similar to 133022 hypotheti	29.68		
45		H20276	Hs.31742	ESTs -	29.24		
	407122 400287	S39329	Hs.181350	kalikrein 2, prostatio	28.90		
	432244	Al669973	Hs.200574	ESTs	28.74		
		U80456	Hs.27311	single-minded (Drosophila) homolog 2	28.74		
50	451939 415989	AI267700	Hs.111128	ESTs	28.34		
30		AW967648	Hs.23023	ESTs	27.34		
	418961 425628	NM 004476	Hs.1915	folate hydrotase (prostate-specific memb	27.32		
			Hs.282906	ESTs	27.24		
	458509 448290	AA654650 AK002107	Hs.20843	Homo sapiens cDNA FLJ11245 fis, clone PL	27.16		
55			Hs.183752	microseminoprotein, beta-	26.17		
55	428336	AA503115		holocarboxylase synthetase (biotin-[prop	25.60		
	450096	Al682088	Hs.223368	kallikrein 3. (prostate specific antigen	24.91		
	400299	X07730	Hs.171995		24.91		
	437571	AA760894	Hs.153023	ESTS	24.74		
60	453160	Al263307	Hs.146228	H2B histone femily, member L	24.66		
oo	453096	AW294631	Hs.11325	ESTs acid phosphatase, prostate	24.40		
	425075	AA506324	Hs.1852		24.23		
	407202	N58172	Hs.109370	ESTs	24.18		

	424846	AU077324	Hs.1832	neuropeptide Y	23.57
	453370	AJ470523	Hs.182356	ATP-binding cassette, sub-family C (CFTR	23.16
	422805	AA436989	Hs.121017	H2A histone family, member A	22.52
~	444917	R68651	Hs.144997	ESTs	22.26
5	408826	AF216077	Hs.48376	Homo sapiens clone HB-2 mRNA sequence	22.02
	413597	AW302885	Hs.117183	ESTs	21.76
	426429	X73114	Hs.169849	myosin-binding protein C, slow-type	21,32
	435981	H74319	Hs.188620	ESTs	21.12
10	432966	AA650114		ESTs	21.07
10	418848	AI820961	Hs.193465	ESTs	21.06
	405685				20.90
	443271	BE568568	Hs.195704	ESTs	19.98
	418819	AA228776	Hs.191721	ESTs	19.94
15	420757	X78592	Hs.99915	androgen receptor (dihydrotestosterone r	19.72
13	418994	AA296520	Hs.89546	selectin E (endothellal adhesion molecul	19.56
	429918	AW873986	Hs.119383	ESTs	19.04
	415539	A1733881	Hs.72472	ESTs	18.43
	450382	AA397658	Hs.60257	Homo sapiens cDNA FLJ13598 fis, clone PL	18,34
20	418829 429984	AA516531	Hs.55999	NK homeobox (Drosophila), family 3, A	18.28
20	443822	AL050102 Al087412	Hs.227209 Hs.143611	hypothetical protein FLJ21617	17.82
				ESTs, Weakly similar to 2004399A chromos	17.66
	431676 410330	AI685464 AW023630	Hs.292638 Hs.46786	gb:tt98f04.x1 NCI_CGAP_Pr28 Homo saplens ESTs	17.84 17.52
	432441	AW023630 AW292425	Hs.163484	FSTs	17.52
25	452792	AB037765	Hs.30652	KIAA1344 protein	17.39
23	445472	AB006631	Hs.12784	Homo saniens mRNA for KIAA0293 gene, par	17.00
	414565	AA502972	Hs.183390	hypothetical protein FLJ13590	16.82
	430487	D87742	Hs.241552	KIAA0268 protein	16.72
	431716	D89053	Hs.268012	fatty-acid-Coenzyme A ligase, long-chain	16.60
30	419536	AA603305	HS-200012		16.50
50	439677	R82331	Hs.164599	gb:np12d11,s1 NCI_CGAP_Pr3 Homo sapiens ESTs	16,46
	449625	NM 014253	Hs.23796	odz (odd Oz/ten-m, Drosophila) homolog 1	16.32
	408430	S79876	Hs.44926	dipeptidylpeptidase IV (CD26, adenosine	16.28
	447033	A/357412	Hs.157601	ESTs	16.02
35	453006	Al362575	Hs.167133	ESTs	15.74
55	431474	AL133990	Hs.190842	ESTs	15.74
	420218	AW958037	Hs.22437	ribosomal protein L4	15.64
	408000	L11690	Hs.620	bullous pemphigoid antigen 1 (230/240kD)	15.54
	416208	AW291168	Hs.41295	ESTs, Weakly similar to MUC2_HUMAN MUCIN	15.48
40	430226	BE245562	Hs.2551	adrenergic, beta-2-, receptor, surface	15.40
40	415263	AA948033	Hs.130853	ESTs	15.38
	432437	W07088	Hs.293665	ESTs	15.26
	428398	Al249368	Hs.98558	ESTs	15.21
	429900	AA460421	Hs.30875	ESTs	14.90
45	449156	AF103907	Hs.171353	prostate cancer antigen 3	14.89
•••	411096	U80034	Hs.68583	mitochondrial Intermediate peptidase	14.81
	435974	U29690	Hs.37744	Homo sapiens beta-1 adrenergic receptor	14.76
	444484	AK002126	Hs.11260	hypothetical protein FLJ11264	14.76
	422728	AW937826	Hs.103262	ESTs, Weakly similar to ZN91_HUMAN ZINC	14.60
50	418601	AA279490	Hs.86368	calmedia	14.56
-	448999	AF179274	Hs.22791	transmembrane protein with EGF-like and	14.55
	445835	AI734009	Hs.127699	KIAA1603 protein	14.44
	452712	AW638616	112111000	gb:RC5-LT0054-140200-013-D01 LT0054 Homo	14.22
	432189	AA527941		gb:nh30c04.s1 NCL CGAP Pr3 Homo sapiens	14.12
55	424565	AW102723	Hs.75295	guanylate cyclase 1, soluble, alpha 3	13.78
	429290	AP203032	Hs.198760	neurofilament, heavy polypeptide (200kD)	13.57
	419264	AA877104	Hs.293672	ESTs, Weakly similar to ALUB_HUMAN !!!!	13.40
	416445	AL043004	Hs.300678	KIAA0135 protein	13.32
	407275	Al364186	110200070	gb:qw34h07.x1 NCI_CGAP_Ut4 Homo sapiens	13.24
60	408369	R38438	Hs.182575	solute carrier family 15 (H+/peptide tra	13.21
	446720	Al439136	Hs.140546	ESTs	13.06
	434988	Al418055	Hs.161160	ESTs	13.02
	448172	N75276	Hs.135904	ESTs	12.98
	416182	NM_004354	Hs.79069	cyclin G2	12.94
65	420544	AA677577	Hs.98732	Homo sapiens Chromosome 16 BAC clone CIT	12.79
	445413	AA151342	Hs.12677	CGI-147 protein	12.64
	452588	AA889120	Hs.110637	homeo box A10	12.62
	407819	R42185	Hs.274803	ESTs	12.60
	433444	AW975324	Hs.129816	ESTs	12.60

	421059	AI854133	Hs.30212	thyroid receptor interacting protein 15	12.30
	420077	AW512260	Hs.87767	ESTs	12.24
	453930	AA419466	Hs.36727	hypothetical prolein FLJ10903	12.22
5	441610	AW576148	Hs.148376	ESTs	12.20
3	451009	AA013140	Hs.115707	ESTs	12.16
	433764 440286	AW753676 U29589	Hs.39982 Hs.7138	ESTs	12.04
	443912	D29069 R37257	Hs.184780	cholinergic receptor, muscarinic 3 ESTs	11.92
	419526	Al821895	Hs.193481	ESTs	11.91
10	423073	BE252922	Hs.123119	MAD (mothers against decapentaplegic, Dr	11.87
10	452784	BE463857	Hs.151258	hypothetical protein FLJ21062	11.88
	414422	AA147224	Hs.71814	ESTs	11.76
	450203	AF097994	Hs.301528	L-kynurenine/alpha-eminoadipate aminotra	11.68
	436679	Al127483	Hs.120451	ESTs, Weakly similar to unnamed protein	11.60
15	440901	AA909358	Hs.128612	ESTs	11.60
	448045	AJ297436	Hs.20166	prostate stem cell antigen	11.51
	433887	AW204232	Hs.279522	ESTs	11.50
	434980	AW770553	Hs.293640	sterol O-acyltransferase (acyl-Coenzyme	11.35
20	425905	AB032959	Hs.161700	novel C3HC4 type Zinc finger (ring finge	11.33
20	434680	T11738	Hs.127574 Hs.297647	ESTs calcium channel, voltage-dependent, L ty	11.18
	449650 431173	AF055575 AW971198	Hs.294068	ESTs	11.16
	434539	AW748078	Hs.214410	ESTs, Weakly similar to MUC2_HUMAN MUCIN	11.16
	410037	AB020725	Hs.58009	KIAA0918 protein	11.14
25	417708	N74392	Hs.50495	ESTs	11.14
	458332	AJ000341	Hs.220491	ESTs	11.12
	420381	D50640	Hs.301782	phosphodiesterase 3B, cGMP-inhibited	11.10
	425665	AK001050	Hs.159066	hypothetical protein FLJ10188	11.08
20	425710	AF030880	Hs.159275	solute carrier family, member 4	11.08
30	428728	NM_016625	Hs.191381	hypothetical protein	11.04
	407021 410733	U52077 D64264	Hs.66052	gb:Human mariner1 transposase gene, comp CD38 antigen (p45)	11.02
	401714	D64264	H8.00002	CDS8 anagen (p4c)	10.90
	434485	Al623511	Hs.118567	ESTs	10.89
35	415786	AW419196	Hs.257924	hypothetical protein FLJ 13782	10.87
	452340	NM_002202	Hs.505	ISL1 transcription factor, LIM/homeodoma	10.85
	453628	AW243307	Hs.170187	hypothetical protein	10.72
	408063	BE086548	Hs.42346	calcineurin-binding protein calsarcin-1	10.67
	417687	AI828596	Hs.250691	ESTs	10.64
40	434666	AF151103	Hs.112259	T cell receptor gamma locus	10.53
	432374	W68815	Hs.301885	Homo sapiens cDNA FLJ11346 fis, clone PL	10.50
	428819 413409	AL135623 Al638418	Hs.193914 Hs.21745	KIAA0575 gene product DEAD/H (Asp-Glu-Ala-Asp/His) box polypep	10.44
	428775	AA434579	Hs.143691	ESTs	10.21
45	436556	Al364997	Hs.7572	ESTs	10.20
	441690	R81733	Hs.33106	ESTs	10.14
	419852	AW503756	Hs.286184	hypothetical protein dJ551D2.5	10.10
	421991	NM_014918	Hs.110488	KIAA0990 protein	10.04
	423698	AA329796	Hs.1098	DKFZp434J1813 protein	10.02
50	452039	Al922988	Hs.172510	ESTs	10.00
	433043	W57554	Hs.125019	ESTs	9.98
	433927	AI557019	Hs.116467	small nuclear protein PRAC	9.97 9.96
	445424	AB028945	Hs.12696	cortactin SH3 domain-binding protein Homo sapiens cDNA FLJ13581 fis, clone PL	9.88
55	432240 433104	Al694767 AL043002	Hs.129179 Hs.128246	ESTs, Moderately similar to unnamed prot	9.84
55	452744	AI267652	Hs.30504	Homo saplens mRNA; cDNA DKFZp434E082 (fr	9.82
	431217	NM 013427	Hs.250830	Rho GTPase activating protein 6	9.75
	427398	AW390020	Hs.20415	chromosome 21 open reading frame 11	9.70
	446896	T15767	Hs.22452	Homo saplens mRNA for KIAA1737 protein,	9.70
60	421470	R27496	Hs.1378	annexin A3	9.64
	406554				9.60
	401424				9.58
	407902	AL117474	Hs.41181	Homo sapiens mRNA; cDNA DKFZp727C191 (fr	9.56
65	423545	AP000692	Hs.129761 Hs.35596	chromosome 21 open reading frame 5 ESTs	9.54 9.51
U.J	439024 431548	R96696 AI834273	Hs.9711	novel protein	9.48
	409262	AK000631	Hs.52256	hypothetical protein FLJ20624	9.45
	446271	D82484	Hs.100469	ESTs	9.42
	449692	AW013907	Hs.224276	methylcrotonoyl-Coenzyme A carboxylase 2	9.26

	414140	AA281279	Hs.23317	hypothetical protein FLJ14681	9.24
	435980	AF274571	Hs.129142	deoxyribonuclease II beta	9.24
	421246	AW582962	Hs.300961	CGI-47 protein	9.20
5	427304	AA761526	Hs.163853	ESTs	9.16 9.16
3	442914 413627	AW188551 BE182082	Hs.99519 Hs.246973	hypothetical protein FLJ14007 ESTs	9.14
	439699	AF086534	Hs.187561	ESTS, Moderately similar to ALU1_HUMAN A	9.10
	437718	Al927288	Hs.196779	ESTs	9.07
	439820	AL360204	Hs.283853	Homo sapiens mRNA full length insert cDN	9.06
10	447342	Al199268	Hs.19322	Homo sapiens, Similar to RIKEN cDNA 2010	9.05
	446223	BE300091	Hs.119699	hypothetical protein FLJ12969	9.04
	410001	AB041036	Hs.57771	kallikrein 11	9.03
	424012	AW368377	Hs.137569	tumor protein 63 kDa with strong homolog	9.03
	441791	AW372449	Hs.175982	hypothetical protein FLJ21159	9.02
15	448206	BE622585	Hs.3731	ESTs, Moderately similar to 138022 hypot	9.02
	414269	AA298489		olfactory receptor, family 51, subfamily	8.99
	442081	AA401863	Hs.22380	ESTs	8.98
	420092	AAB14043	Hs.88045	ESTs	8.85
20	411630 421863	U42349 AI952677	Hs.71119 Hs.108972	Putative prostate cancer turnor suppresso Homo sapiens mRNA; cDNA DKFZp434P228 (fr	8.80 8.80
20	454141	AW138413	Hs.182356	ATP-binding cassette, sub-family C (CFTR	8.80
	418278	AI088489	Hs.83937	hypothetical protein	8.78
	428330	L22524	Hs.2256	matrix metalloproteinase 7 (matrilysin,	8.76
	432415	T16971	Hs.289014	ESTs, Weakly similar to A43932 mucin 2 p	8.75
25	424906	AI566086	Hs.153716	Homo sapiens mRNA for Hmob33 protein, 3'	8.74
	415245	N59650	Hs.27252	ESTs	8.72
	442409	BE208843	Hs.129544	hypothetical protein MGC15438	8.70
	404571			" '	8.66
	418033	W68180	Hs.259855	elongation factor-2 kinase	8.64
30	456497	AW967956	Hs.123648	ESTs, Weakly similar to AF108460 1 ubinu	8.56
	405876				8.54
	448807	AI571940	Hs.7549	ESTs	8.52
	445372	N36417	Hs.144928 Hs.300615	ESTs ESTs	8.48 8.44
35	425171	AW732240 X04430	Hs.93913		8.36
33	419968 407385	AA610150	Hs.272072	interleukin 6 (interferon, beta 2) ESTs, Wealdy similar to I36022 hypotheti	8.31
	433172	AB037841	Hs.102652	hypothetical protein ASH1	8.30
	422631	BE218919	Hs.118793	hypothetical protein FLJ10688	8.27
	412719	AW016610	Hs.129911	ESTs	8.24
40	418849	AW474547	Hs.53565	Homo sapiens PIG-M mRNA for mannosvitran	8.22
	444922	Al921750	Hs.144871	Homo sapiens cDNA FLJ13752 fis, clone PL	8.22
	427674	NM_003528	Hs.2178	H2B histone family, member Q	8.20
	432101	Al918950	Hs.11092	EphA3	8.17
45	416288	H51299		gb:yp07c03.s1 Scares breast 3NbHBst Homo	8.15
45	404915			10111111	8.08
	440106	AA864988	Hs.127699 Hs.57787	KIAA1603 protein ESTs	8.07 8.06
	442861 452259	AA243837 AA317439	Hs.28707		8.06
	443250	AI041530	Hs.132107	signal sequence receptor, gamma (translo ESTs	8.06
50	437267	AW511443	Hs.258110	ESTs	8.04
50	452891	N75582	Hs.212875	ESTs, Weakly similar to DYH9_HUMAN CILI	8.02
	422219	AW978073		regulator of mitotic spindle assembly 1	8.00
	453049	BE537217	Hs.30343	ESTs	8.00
	439731	Al953135	Hs.45140	hypothetical protein FLJ14084	7.98
55	408554	AA836381	Hs.7323	nuclear receptor co-repressor/HDAC3 comp	7.94
	421154	AA284333	Hs.287631	Homo sapiens cDNA FLJ14269 fis, clone PL	7.94
	430107	AA465293	Hs.105069	ESTs	7.94
	433404	T32982	Hs.102720	ESTs	7.93
60	450813	Al739625	Hs.203376	ESTs	7.90
00	416239 448212	AL038450 Al475858	Hs.48948	ESTs gb:tc87d07.x1 NCL_CGAP_CLL1 Homo saplens	7.85 7.82
	448212 449532	W74653	Hs.271593	ESTs, Moderately similar to A47532 B-cel	7.82
	413930	W/4003 M86153	Hs.75618	RAB11A, member RAS oncogene family	7.80
	458191	Al420611	Hs.127832	ESTs	7.80
65	444858	Al199738	Hs.208275	ESTs, Weakly similar to ALUA_HUMAN !!!!	7.78
	457498	Al732230	Hs.191737	ESTs	7.78
	407235	D20569	Hs.169407	SAC2 (suppressor of actin mutations 2, y	7.76
	433759	AA680003	Hs.109363	Homo saplens cDNA: FLJ23603 fis, clone L	7.74
	433805	AA706910	Hs.112742	ESTs	7.74

7.72

	426485	NM_000207	HS.170040	bigreier-geuseg Browitt recipi recebror-	1.12
	446028	R44714	Hs.106795	Homo sapiens cDNA FLJ13136 fis, clone NT	7.72
	418555	Al417215	Hs.87159	hypothetical protein FLJ12577	7.70
	447499	AW262580	Hs.147674	protocacherin beta 16	7.70
5	419839	U24577	Hs.93304	phospholipase A2, group VII (platelet-ac	7.68
,					7.68
	416857	AA188775	Hs,292453	ESTs	
	413801	M62246	Hs.35406	ESTs, Highly similar to unnamed protein	7.66
	425480	AB023198	Hs.158135	KIAA0981 protein	7.66
	420120	AL049610	Hs.95243	transcription elongation factor A (SII)-	7.64
10	424099	AF071202	Hs.139336	ATP-binding cassette, sub-family C (CFTR	7.64
10					7.63
	446307	T50083	Hs.9094	ESTs	
	429220	AW207208	Hs.136319	ESTs	7.59
	420345	AW295230	Hs.25231	ESTs	7.54
	429208	AA447990	Hs.190478	EŜTs	7.54
15	447247	AW369351	Hs.267955	Homo sapiens cDNA FLJ13090 fis, clone NT	7.53
13			Hs.10263	ESTs	7.53
	440995	T57773			7.52
	448706	AW291095	Hs.21814	interleukin 20 receptor, alpha	
	410227	AB009284	Hs.61152	exostoses (multiple)-like 2	7.49
	431616	AA508552	Hs.195839	ESTs, Weakly similar to 138022 hypotheti	7.46
20	434217	AW014795	Hs.23349	ESTs	7.44
20	431467	N71831	Hs.256398	Homo sapiens mRNA; cDNA DKFZp434E0528 (f	7.42
			Hs.244334	Homo sapiens prostein mRNA, complete cds	7.42
	448519	AW175665			7.40
	448791	Al632278	Hs.34981	ESTs	
	419743	AW408762	Hs.127478	Homo sapiens clone 24416 mRNA sequence	7.39
25	445855	BE247129	Hs.145569	ESTs	7.36
	425211	M18667	Hs.1867	progastricsin (pepsinogen C)	7.35
	419131	AA406293	Hs.301622	ESTs	7.34
					7,33
	400294	N95796	Hs.179809	Homo sapiens prostein mRNA, complete cds	7.28
	441736	AW292779	Hs.169799	ESTs	
30	427701	AA411101	Hs.221750	nuclear autoantigenic sperm protein (his	7.24
	457733	AW974812	Hs.291971	ESTs	7.24
	418432	M14156	Hs.85112	insulin-like growth factor 1 (somatomedi	7.22
	441201	AW1 18822	Hs.128757	ESTs	7.21
			Hs.125752	ESTs	7.20
25	419953	BE267154			7.20
35	419991	AJ000098	Hs.94210	eyes ebsent (Drosophila) homolog 1	
	425018	BE245277	Hs.154196	E4F transcription fector 1	7.20
	424560	AA158727	Hs.150555	protein predicted by clone 23733	7.18
	435380	AA679001	Hs.192221	ESTs	7.14
	420658	AW965215	Hs.130707	ESTs	7.12
40				KIAA0974 protein	7.10
40	408291	AB023191	Hs.44131		
	409110	AA191493	Hs.48778	niban protein	7.10
	414485	W27028	Hs.182625	VAMP (veside-associeted membrane protei	7.10
	430039	BE253012	Hs.153400	ESTs, Weakly similar to ALU1_HUMAN ALU S	7.10
	450832	AW970602	Hs.105421	ESTs	7.10
45	417153	X57010	Hs.81343	collagen, type II, alpha 1 (primary oste	7.08
7.7				ESTs	7.07
	412446	Al768015	Hs.92127		7.06
	412953	Z45794	Hs.238809	ESTs	
	418051	AW192535	Hs.19479	ESTs	7.06
	421566	NM_000399	Hs.1395	early growth response 2 (Krox-20 (Drosop	7.04
50	446999	AA151520	Hs.279525	hypothetical protein MGC4485	7.04
	440529	AW207640	Hs.16478	Homo sapiens cDNA: FLJ21718 fis, clone C	7.04
			Hs.126594	ESTs	7.01
	441111	AI806867			7.00
	451027	AW519204	Hs.40808	ESTs -	
	408432	AW195262		gb:xn67b05.x1 NCI_CGAP_CML1 Homo sapiens	7.00
55	432223	AA333283	Hs.285336	Homo sapiens, clone IMAGE:3460280, mRNA	7.00
	444805	AB007899	Hs.12017	homolog of yeast ubiquitin protein ligas	6.99
	414212	AA136569	Hs.295940	KIAA0187 gene product	6.98
			Hs.2839		6.98
	431725	X65724		Norrie disease (pseudoglioma)	
	449685	AW296669	Hs.66095	ESTs	6.97
60	447313	U92981	Hs.18081	Homo saplens clone DT1P1B6 mRNA, CAG rep	6.98
	424590	AW966399	Hs.46821	hypothetical protein FLJ20086	6.94
	449655	AI021987	Hs.59970	ESTs	6.92
	419563	AA526235	Hs.193162	Homo sapiens cDNA FLJ11983 fis, clone HE	6.90
					6.89
	434163	AW974720	Hs.25206	group XII secreted phospholipase A2	
65	415809	Z32789	Hs.46601	ESTs	6.86
	425782	U66468	Hs.159525	cell growth regulatory with EF-hand doma	6.85
	417958	AA767382	Hs.193417	ESTs	8.84
	427408	AA583206	Hs.2156	RAR-related orphen receptor A	6.79
	445873	AA250970	Hs.251946	poly(A)-binding protein, cytoplasmic 1-l	6.74
	440070	MM230370	110201040	berita a menta brown a alrebitation to	3.7

426485 NM\_006207 Hs.170040 platelet-derived growth fector receptor-

	410718 432383	Al920783 AA534489	Hs.191435	ESTs gb:nf76g11.s1 NCL_OGAP_Co3 Homo sapiens	6.7
	436521	AW203986	Hs.213003	ESTs	6.7
	435604	AA625279	Hs.26892	uncharacterized bone marrow protein BM04	6.7
5	419083	Al479560	Hs.98613	Homo sapiens cDNA FLJ12292 fis, clone MA	6.7
-	418245	AA088767	Hs.83883	transmembrane, prostate androgen induced	6.7
	420714	BF172704	Hs.222746	KIAA1610 protein	6.7
	412707	AW206373	Hs.16443	Homo sapiens cDNA: FLJ21721 fis, clone C	6.6
	421898	N62293	Hs.45107	ESTs	6.6
0	411078	Al222020	Hs.182364	CocoaCrisp	6.6
-	452485	AA810211	Hs.34244	ESTs	6.6
	422763	AA033699	Hs.83938	ESTs, Moderately similar to MAS2_HUMAN M	6.6
	444618	AV653785	Hs.300171	ELL-RELATED RNA POLYMERASE II, ELONGATIO	6.6
	450164	Al239923	Hs.30098	ESTs	6.6
.5	431060	AF039307	Hs.249171	homeo box A11	6.6
	408031	AA081395	Hs.42173	Homo sapiens cDNA FLJ10366 fis, clone NT	6.6
	420285	AA258124	Hs.293878	ESTs, Moderately similar to ZN91_HUMAN Z	6.6
	444670	H58373	Hs.37494	hypothetical protein MGC5370	6.6
	444489	Al151010	Hs.157774	ESTs	6.6
0	445685	AW779829	Hs.263436	gb:hn88a05.x1 NCI_CGAP_Kid11 Homo sapien	6.6
	435677	AA694142	Hs.293726	ESTs, Weakly similar to TSGA RAT TESTIS	6.5
	452221	C21322	Hs.11577	hypothetical protein FLJ22242	6.5
	431510	AA580082	Hs.112264	ESTs	8.5
	415874	AF091622	Hs.78893	KIAA0244 protein	6.5
5	418405	AI868282	Hs.11898	ESTs, Highly similar to KIAA1370 protein	6.5
	452768	AW069459	Hs.61539	ESTs	6.5
	401451				6.5
	416289	W26333		ESTs	6.5
	431778	AL000276	Hs.268562	regulator of G-protein signalling 17	6.5
0	409089	NM_014781	Hs.50421	KIAA0203 gene product	6.5
	442833	AA328153	Hs.88201	ESTs, Weakly similar to A Chain A, Cryst	6.5
	431992	NM_002742	Hs.2891	protein kinase C, mu	6.4
	418833	AW974899	Hs.292776	ESTs	8.4
	429163	AA884766		gb:am20a10.s1 Soares_NFL_T_GBC_S1 Homo s	6.4
35	430403	AF039390	Hs.241382	tumor necrosis factor (ligand) superfami	6.4
	443058	AW451642	Hs.16732	ESTs	6.4
	418564	AA631143	Hs.179809	Homo sapiens prostein mRNA, complete cds	6.4
	432674	AA641092	Hs.257339	ESTs, Weakly similar to I38022 hypotheti	6.4
	423600	Al633559	Hs.29076	ESTs	6.4
ю	404253				6.4
	433610	AA806822	Hs.112547	ESTs	6.4
	421552	AF026692	Hs.105700	secreted frizzled-related protein 4	6.4
	407118	AA156790	Hs.262036	ESTs, Weakly similar to Z223_HUMAN ZINC	6.4
	408606	N79738	Hs.136102	KIAA0853 protein	6.4
5	421452	Al925946	Hs.104530	fetal hypothetical protein	6.4
	433285	AW975944	Hs.237396	ESTs	6.4
	434926	BE543269	Hs.50252 Hs.214013	mitochondrial ribosomal protein L32 FSTs	6.4
	446189	H85224			6.3
60	416806	NM_000288	Hs.79993	peroxisomal biogenesis factor 7	6.3
v	416467	H57585	Hs.61779	ESTS Home conjune cOMA EL H3501 fis close PI	6.3
	453403	BE466639	Hs.218366	Homo sapiens cDNA FLJ13591 fis, clone PL	6.3
	429769	NM_004917	Hs.157148	kallikrein 4 (prostase, enamel matrix, p hypothetical protein MGC13204	6.3
	423642	AW452650 BE313280	Hs.15/148 Hs.159627		6.3
55	425843 439221	AA737106	Hs.32250	death associated protein 3 ESTs, Moderately similar to 178885 serin	6.1
J	439221 428194	AA787106 AA765603	Hs.180877	H3 histone, family 3B (H3.3B)	8.3
	428194	X63629	Hs.2877	cadherin 3, type 1, P-cadherin (placanta	6.3
	431958	AF100143	Hs.6540	fibroblast growth factor 13	6.1
	452789	AW081626	Hs.242561	ESTs	6.
60	452789	D54745	Hs.80247	cholecystokinin	6.5
,,,	436962	AW377314	Hs.5364	DKFZP564I052 protein	6.1
	433383	AF034837	Hs.192731	double-stranded RNA specific adenosine d	6.3
	433383	AW749855	110.182731	gb:QV4-BT0534-281299-053-c05 BT0534 Homo	6.1
	450728	AW162923	Hs.25363	preseniin 2 (Alzheimer disease 4)	6.1
55	440293	AID04193	Hs.22123	ESTs	6.3
,,	453745	AA952969	Hs.63908	hypothetical protein MGC14726	6.3
	426595	AW971980	Hs.82402	p21/Cdc42/Rac1-activated kinase 1 (yeast	6.3
	444412	Al147652	Hs.216381	Homo saplens clone HH409 unknown mRNA	6.3
	413384	NM_000401	Hs.75334	exostoses (multiple) 2	6.2

	426320 423349	W47595 AF010258	Hs.169300 Hs.127428	transforming growth factor, beta 2 homeo box A9	6.22
	429165	AW009888	Hs.118258	prostate cancer associated protein 1	6.18
	424800	AL035588	Hs.153203	MyoD family inhibitor	6.18
5	409564	AA045857	Hs.54943	fracture callus 1 (rat) homolog	6.16
-	438796	W67821	Hs.109590	genethonin 1	6.16
	425451	AF242769	Hs.157461	mesenchymal stem cell protein DSC54	8.14
	451663	AI872360	Hs.209293	ESTs	6.14
	413623	AAB25721	Hs.246973	ESTs	8.12
10	452232	AW020603	Hs.271698	radial spoke protein 3	6.12
	453390	AAB62496	Hs.28482	ESTs	6.12
	435542	AA687376	Hs.269533	ESTs	6.12
	420424	AB033036	Hs.97594	KIAA1210 protein	6.11
10	407103	AA424881	Hs.256301	hypothetical protein MGC13170	6.10
15	409734	BE161664	Hs.56155	hypothetical protein	6.10 6.10
	432686	BE223007	Hs.152460	Homo sapiens cDNA FLJ12909 fis, clone NT Homo sapiens cDNA: FLJ23077 fis, clone L	6.10
	438361 411479	AA805666 AW848047	Hs.148217	gb:iL3-CT0214-291299-052-A12 CT0214 Homo	8.10
	438849	W28948	Hs.10762	ESTs	6.08
20	452726	AF188527	Hs.61661	ESTs. Weakly similar to AF174605 1 F-box	6.08
20	445895	D29954	Hs.13421	KIAA0056 protein	6.08
	440774	Al420611	Hs.127832	ESTs	6.07
	422583	AA410506	Hs.118578	KIAA0874 protein	6.06
	427500	AW970017	Hs.293948	ESTs, Weakly similar to S65657 alpha-1C-	6.04
25	443646	AI085198	Hs.298699	ESTs	6.04
	410566	AA373210	Hs.43047	Homo sapiens cDNA FLJ13585 fis, clone PL	6.02
	417845	AL117461	Hs.82719	Homo saplens mRNA; cDNA DKFZp586F1822 (f	8.02
	430273	Al311127	Hs.125522	ESTs	6.02
••	434792	AA649253	Hs.132458	ESTs	6.01
30	442490	AW965078	Hs.30212	thyroid receptor interacting protein 15	6.01
	420026	Al831190	Hs.166678	ESTs	6.00 6.00
	437782	AI370876	Hs.123163	exportin 1 (CRM1, yeast, homolog) adenylate kinase 5	6.00
	447359 447713	NM_012093 Al420733	Hs.18268 Hs.207083	ESTs	6.00
35	451073	AI758905	Hs.206063	ESTs	6.00
22	451640	AA195601	Hs.26771	Human DNA sequence from clone 747H23 on	6.00
	410889	X91662	Hs.66744	twist (Drosophila) homolog (acrocephalos	5.97
	441222	Al277237	Hs.44208	hypothetical protein FLJ23153	5.96
	447732	Al758398	Hs.161318	ESTs	5.96
40	437756	AA767537	Hs.197096	ESTs	5.95
	406829	NM_006042	Hs.48384	heparan sulfate (glucosamine) 3-O-sulfot	5.94
	453911	AW503857	Hs.4007	Sarcolemmal-associated protein	5.94
	414085	AA114016	Hs.75746	aldehyde dehydrogenase 1 family, member	5.93
40	406875	NM_015434	Hs.48604	DKFZP434B168 protein	5.92
45	439451	AF086270	Hs.278554	heterochromatin-like protein 1	5.92 5.91
	423853	AB011537 AW294092	Hs.133466 Hs.21594	siit (Drosophila) homolog 1 hypothetical protein MGC15754	5.91
	453060 420407	AA814732	Hs.145010	lipopolysaccaride-specific response 5-li	5.91
	450480	X82125	Hs.25040	zinc finger protein 239	5.90
50	408446	AW450669	Hs.45068	hypothetical protein DKFZp434I143	5.88
	421039	NM 003478	Hs.101299	cullin 5	5.88
	451684	AF216751	Hs.26813	CDA14	5.88
	436063	AK000028	Hs.250867	ribosomal protein S24 -	5.86
	410507	AA355288	Hs271408	transitional epithelia response protein	5.86
55	420179	N74530	Hs.21168	ESTs	5.84
	453878	AW964440	Hs.19025	DC32	5.84
	452270	AW975014	Hs.26	ferrochelatase (protoporphyria)	5.83
	435887	AA954229	Hs.114052 Hs.239154	ESTs	5.82 5.82
60	417683	AW566008		ankyrin repeat, family A (RFXANK-like), ESTs, Weakly similar to ELL2_HUMAN RNA P	5.81
JU	432005 406815	AA524190 AA833930	Hs.120777 Hs.288096	tRNA isopentenylpyrophosphate transferas	5.80
	437980	R50393	Hs.278436	KIAA1474 protein	5.80
	425856	AA364908	Hs.98927	hypothetical protein FLJ13993	5.79
	400301	X03635	Hs.1657	estrogen receptor 1	5.78
65	446261	AA313893	Hs.13399	hypothetical protein FLJ12615 similar to	5.78
	410141	R07775	Hs.287657	Homo sapiens cDNA: FLJ21291 fis, clone C	5.77
	427258	AA400091	Hs.39421	ESTs	5.76
	419108	AA389724	Hs.191264	ESTs, Weakly similar to ALU7_HUMAN ALU S	5.76
	442029	AW956698	Hs.14456	neural precursor cell expressed, develop	5.76

	407783	AW996872	Hs.172028	a disintegrin and metalloproteinase doma	5.75
	434408	AI031771	Hs.132586	ESTs	5.74
	415077	L41607	Hs.934	glucosaminyl (N-acetyl) transferase 2, I	5.74
-	432435	BE218886	Hs.282070	ESTs	5.74
5	433313	W20128 N75450	Hs.296039 Hs.183412	ESTs	5.73 5.73
	431740 412991	AW949013	HS.103412	ESTs, Moderately similar to AF118721 67 gb:QV4-FT0005-110500-201-e12 FT0005 Homo	5.72
	418852	BE537037	Hs.273294	hypothetical protein FLJ20069	5.72
	418882	NM 004996	Hs.89433	ATP-binding cassette, sub-family C (CFTR	5.72
10	446867	AB007891	Hs.16349	KIAA0431 protein	5.72
	437866	AA156781	Hs.83992	metallothionein 1E (functional)	5.72
	410232	AW372451	Hs.61184	CGI-79 protein	5.70
	414452	AA454038	Hs.29032	ESTs	5.70
	422762	AL031320	Hs.119976	Human DNA sequence from clone RP1-20N2 o	5.70
15	428730	AA825947	Hs.25750	ESTs	5.70
	431571	AW500486	Hs.180610	splicing factor proline/glutamine rich (	5.70
	433393	AF038564 AL133067	Hs.98074 Hs.25214	itchy (mouse homolog) E3 ubiquitin prote hypothetical protein	5.70 5.70
	450616 443774	AL133007 AL117428	Hs.9740	DKFZP434A236 protein	5.69
20	446100	AW967109	Hs.13804	hypothetical protein dJ462O23.2	5.69
20	419168	AI336132	Hs.33718	Homo sapiens cDNA FLJ12641 fis, clone NT	5,68
	416653	AA768553	Hs.77496	metallothionein 1E (functional)	5.67
	452679	Z42387	Hs.4299	transmembrane, prostate androgen induced	5.66
	450244	AA007534	Hs.125062	ESTs	5.66
25	408621	AJ970672	Hs.46638	chromosome 11 open reading frame 8	5.65
	450325	AI935962	Hs.26289	ESTs	5.65
	439671	AW162840	Hs.6641	kinesin family member 5C	5.64 5.64
	452387 413992	AI680772 W26276	Hs.4316 Hs.136075	trinucleotide repeat containing 12 RNA, U2 small nuclear	5.63
30	444151	AW972917	Hs.128749	alpha-methylacyl-CoA racemase	5.63
50	417791	AW965339	Hs.111471	ESTs	5.62
	410196	Al936442	Hs.59838	hypothetical protein FLJ10908	5.60
	415123	D60925		ESTs	5.60
~~	429170	NM_001394	Hs.2359	dual specificity phosphatase 4	5.60
35	434415	BE177494		gb:RC6-HT0596-270300-011-C05 HT0596 Homo	5.60
	440738 443830	AI004650 AI142095	Hs.225674 Hs.143273	WD repeat domain 9 ESTs	5.60 5.60
	449603	A1142095 A1655662	Hs.197698	ESTS	5.60
	414342	AA742181	Hs.75912	KIAA0257 protein	5.69
40	422634	NM 016010	Hs.118821	CGI-62 protein	5.56
	435047	AA454985	Hs.54973	cadherin-like protein VR20	5.55
	400268				5.55
	452055	Al377431	Hs.293772	hypothetical protein MGC10858	5.54
45	437073	AI885608	Hs.94122	ESTs	5.54
45	434072	H70854	Hs.283059 Hs.104215	Homo sapiens PRO1082 mRNA, complete cds ESTs, Moderately similar to SPCN_HUMAN S	5.53 5.52
	418339 434551	AA639902 BE367162	Hs.280858	ESTs, Highly similar to A35661 DNA excis	5.52
	439569	AW602166	Hs.222399	CEGP1 protein	5.51
	441102	AA973905	Hs.16003	intermediate filement protein syncoilin	5.50
50	448310	AI480316		gb:tm26h09.x1 Soares_NFL_T_GBC_S1 Homo s	5.50
	413173	BE076926	Hs.70980	ESTs	5.48
	436246	AW450963	Hs.119991	ESTs	5.48
	449300	AI656959	Hs.222165	ESTs -	5.48
55	452823 451403	AB012124 AA885569	Hs.30696 Hs.15727	transcription factor-like 5 (basic helix Homo sapiens cDNA FLJ14511 fis, clone NT	5.48 5.46
33	417061	AI675944	Hs.188691	Homo saplens cDNA FLJ12033 fis, done HE	5.44
	429126	AW172356	Hs.99083	ESTs	5.44
	431316	AA502663	Hs.145037	ESTs	5.44
	439192	AW970536	Hs.105413	ESTs	5.44
60	431938	AA938471	Hs.115242	specific granule protein (28 kDa); cysle	5.44
	451552	AA047233	Hs.33810	ESTs	5.43
	416991	N36389	Hs.295091	KIAA0226 gene product	5.42 5.42
	427638 427718	AA406411 Al798680	Hs.208341 Hs.25933	ESTs, Weakly similar to KIAA0969 protein ESTs	5.42
65	438710	AA833907	Hs.178724	ESTs, Weakly similar to ALU1_HUMAN ALU S	5.42
03	406076	AL390179	Hs.137011	Homo sapiens mRNA; cDNA DKFZp547P134 (fr	5.40
	431263	AW129203	Hs.13743	ESTs	5.40
	421264	AL039123	Hs.103042	microtubule-associated protein 1B	5.38
	421685	AF189723	Hs.106778	ATPase, Ca++ transporting, type 2C, memb	5.37

	408460	AA054726	Hs.285574	ESTs	5.36
	409091	AW970386	Hs.269423	ESTs	5.36
	421987	Al133161	Hs.286131	CGI-101 protein	5.36
5	428002	AA418703	11 040040	gb:zv98c03.s1 Soares_NhHMPu_S1 Homo sapi	5.36 5.36
,	441217 426006	AI922183 R49031	Hs.213246 Hs.22627	ESTs ESTs	5.35
	422806	BE314767	Hs.1581	glutathione S-transferase theta 2	5.34
	432281	AK001239	Hs.274263	hypothetical protein FLJ 10377	5.32
	451982	F13036	Hs.27373	Homo sapiens mRNA; cDNA DKFZp564O1763 (f	5.32
10	421129	BE439899	Hs.89271	ESTs	5.31
	444042	NM_004915	Hs.10237	ATP-binding cassette, sub-family G (WHIT	5.31
	410150	AW382942	Hs.6774	ESTs	5.30
	423952	AW877787	Hs.136102	KIAA0853 protein	5.30
	452822	X85689	Hs.288617	hypothetical protein FLJ22621	5.30
15	447752	M73700	Hs.347	factotransferrin	5.29
	441766	R53790	Hs.23294	hypothetical protein FLJ14393	5.29
	431359	AW993522	Hs.292934	ESTs	5.27
	427212	AW293849	Hs.58279	ESTs, Weakly similar to ALU7_HUMAN ALU S	5.27
20	449916	T60525	Hs.299221	pyruvate dehydrogenase kinase, iscenzyme	5.27
20	454014	AW016670	Hs.233275	ESTs	5.27
	419714	AA758751	Hs.98216	ESTs	5.26
	428845	AL157579	Hs.153610	KIAA0751 gene product	5.28 5.24
	417333 419986	AL157545 Al345455	Hs.42179 Hs.78915	bromodomain and PHD finger containing, 3 GA-binding protein transcription factor,	5.24
25	407182	AA312551	Hs.230157	ESTs	5.22
23	420111	AA255652	na.200107	gb:zs21h11,r1 NCl_CGAP_GCB1 Homo sapiens	5.22
	428058	AI821625	Hs.191602	ESTs	5.22
	459551	AI472808	101101000	gb:ti70e07.x1 Soares_NSF_F8_9W_OT_PA_P_S	5.22
	432524	AI458020	Hs.293287	ESTs	5.22
30	436207	AA334774	Hs.12845	hypothetical protein MGC13159	5.22
	410870	U81599	Hs.66731	homeo box B13	5.22
	451418	BE387790	Hs.26369	hypothetical protein FLJ20287	5.22
	409757	NM_001898	Hs.123114	cystatin SN	5.21
~-	441124	T97717	Hs.119563	ESTs	5.21
35	428593	AW207440	Hs.185973	degenerative spermatocyte (homolog Droso	5.21
	436401	AI087958	Hs.29088	ESTs	5.20 5.20
	437113	AA744693	Hs.204662	gb:ny26c10.s1 NCI_CGAP_GCB1 Homo sapiens	5.20
	450947 453279	AI745400 AW893940	Hs.204062 Hs.59698	ESTs ESTs	5.20
40	445467	Al239832	Hs.15617	ESTs, Weakly similar to ALU4_HUMAN ALU S	5.19
40	448944	AB014605	Hs.22599	atrophin-1 interacting protein 1; activi	5.19
	412198	AA937111	Hs.69165	ESTs	5.18
	422646	H87863	Hs.151380	ESTs, Weakly similar to T16584 hypotheti	5.18
	438986	AF085888	Hs.269307	ESTs	5.18
45	453954	AW118338	Hs.75251	DEAD/H (Asp-Glu-Ala-Asp/His) box binding	5.18
	447541	AK000288	Hs.18800	hypothetical protein FLJ20281	5.18
	434029	AA621763	Hs.170434	Homo saplens cDNA FLJ14242 fis, clone OV	5.16
	459294	AW977286	Hs.169531	RBP1-like protein	5.16
EΛ	429441	AJ224172	Hs.204096	lipophilin B (uteroglobin family member)	5.16
50	424692	AA429834	Hs.151791 Hs.79881	KIAA0092 gene product	5.15 5.15
	427359	AW020782	Hs.79881 Hs.146162	Homo sapiens cDNA: FLJ23006 fis, clone L ESTs	5.15
	419872 429422	Al422951 AK001494	Hs.202596	Homo sapiens cDNA FLJ10632 fis, cloné NT	5.14
	448902	Z45998	Hs.22543	Homo sapiens mRNA; cDNA DKFZp76111912 (f	5.14
55	459055	N23235	Hs.30567	ESTs, Weakly similar to B34087 hypotheti	5.14
	431318	AA502700	Hs.293147	ESTs, Moderately similar to A45010 X-lin	5.14
	452953	AJ932884	Hs.271741	ESTs, Weakly similar to A46010 X-linked	5.13
	428372	AK000684	Hs. 183987	hypothetical protein FLJ22104	5.12
	434401	AI864131	Hs.71119	Putative prostate cancer tumor suppresso	5.12
60	416434	AW163045	Hs.79334	nuclear factor, interleukin 3 regulated	5.11
	410268	AA316181	Hs.61635	six transmembrane epithelial antigen of	5.10
	417517	AF001176	Hs.82238	POP4 (processing of precursor, S. cerev	5.10
	453616	NM_003462	Hs.33846	dynein, axonemal, light intermediate pol	5.10
65	427958	AA418000	Hs.98280 Hs.606	potassium intermediate/small conductance	5.09 5.08
05	407945 425154	X69208 NM 001851	Hs.154850	ATPase, Cu++ transporting, alpha polypep collagen, type IX, alpha 1	5.08
	412863	AA121673	Hs.59757	zinc finger protein 281	5.06
	420807	AA280627	Hs.57846	ESTs	5.06
	430568	AA769221	Hs.270847	delta-tubulin	5.06
	-100000				

	433687	AA743991		gb:ny57g01.s1 NCI_CGAP_Pr18 Homo sapiens	5.06
	438375	AW/015940	Hs.232234	ESTs	5.06
	418092	R45154	Hs.106604	ESTs	5.06
5	418576	AW968159	Hs.289104	Alu-binding protein with zinc finger dom	5.05
3	413328	Y15723	Hs.75295	guanylate cyclase 1, soluble, alpha 3	5.04 5.04
	414271 432729	AK000275 AK000292	Hs.75871 Hs.278732	protein kinase C binding protein 1 hypothetical protein FLI20285	5.04
	433433	Al692623	Hs.121513	Homo sapiens clone Z'3-1 placenta expres	5.04
	439662	H97552	Hs.269060	ESTs	5.04
10	439743	AL389956	Hs.283858	Homo sapiens mRNA full length insort cDN	5.04
10	417511	AL049176	Hs.82223	chordin-like	5.02
	437814	AI088192	Hs.135474	ESTs, Weakly similar to DDX9_HUMAN ATP-D	5.02
	426342	AF093419	Hs.169378	multiple PDZ domain protein	5.02
	429782	NM_005754	Hs.220689	Ras-GTPase-activating protein SH3-domain	5.02
15	429975	Al167145	Hs.165538	ESTs	5.02
	436209	AW850417	Hs.254020	ESTs, Moderately similar to unnamed prot	5.02
	438571	AW020775	Hs.56022	ESTs	5.02
	450223	AA418204	Hs.241493	natural killer-tumor recognition sequenc	5.02
20	408267	AW380525	Hs.267705	tubulin-specific chaperone e	5.01 5.00
20	417730	Z44761	11. 4004	gb:HSC28F061 normalized infant brain cDN	5.00
	425465	L18964	Hs.1904	protein kinase C, iota	5.00
	430599 450961	NM_004855 AW978813	Hs.247118 Hs.250867	phosphatidylinositol glycan, class B metallothionein 1E (functional)	5.00
	451388	AR029006	Hs.26334	spestic paraplegia 4 (autosomal dominant	5.00
25	420380	AA640891	Hs.102406	ESTs	4.99
23	424947	R77952	Hs.239825	ESTs. Weakly similar to alternatively so	4.99
	442653	BE269247	Hs.170226	ab:601185486F1 NIH MGC 8 Homo sapiens cD	4.98
	457211	AW972565	Hs.32399	ESTs, Weakly similar to S51797 vasodilat	4.97
	425851	NM_001490	Hs.159642	glucosaminyi (N-acetyl) transferase 1, c	4.97
30	446279	AA490770	Hs.182382	ESTs	4.96
	433377	AI752713	Hs.43845	ESTs	4.96
	450218	R02018	Hs.168640	ankylosis, progressive (mouse) homolog	4.96
	412715	NM_000947	Hs.74519	primase, polypeptide 2A (58kD)	4.94
25	448164	R61680	Hs.26904	ESTs, Moderately similar to Z195_HUMAN Z	4.94
35	420121	AW968271	Hs.191534	ESTs, Weakly similar to ALU1_HUMAN ALU S	4.94 4.93
	421689	N87820 AV655234	Hs.106828 Hs.298083	KIAA1896 protein ESTs, Moderately similar to PC4259 ferri	4.92
	445808 416533	BE244053	Hs.79362	retinoblastoma-like 2 (p130)	4.92
	418049	AA211467	Hs.190488	Homo sapiens, Similar to nuclear localiz	4.92
40	436039	AW023323	Hs.121070	ESTs	4.92
	432653	N62096	Hs.293185	ESTs, Weakly similar to JC7328 amino aci	4.91
	420324	AF163474	Hs.96744	prostate androgen-regulated transcript 1	4.91
	403047				4.91
	436899	AA764852	Hs.291587	ESTs	4.90
45	431117	AF003522	Hs.250500	delta (Drosophila)-like 1	4.90
	427617	D42063	Hs.179825	RAN binding protein 2	4.88
	428804	AK000713	Hs.193738	hypothetical protein FLJ20706	4.88
	433050	AI093930	Hs.163440	Homo sapiens cDNA: FLJ21000 fis, clone C	4.88
50	418575	AA225313	Hs.222888	ESTs, Weakly similar to TRHY_HUMAN TRICH	4.88 4.86
50	432615 412652	AA557191 AI801777	Hs.55028 Hs.6774	ESTs, Weakly similar to I54374 gene NF2 ESTs	4.86
	432473	Al202703	Hs.152414	FSTs	4.86
	449071	NM_005872	Hs.22960	breast carcinoma amplified sequence 2 -	4.86
	450654	AJ245587	Hs.25275	Kruppel-type zinc finger protein	4.85
55	418866	T65754	Hs.100469	gb:yc11c07.s1 Stratagene lung (937210) H	4.85
55	407596	R86913		gb:yg30f05.r1 Soares fetal liver spleen	4.84
	456516	BE172704	Hs.222746	KIAA1610 protein	4.84
	426501	AW043782	Hs.293818	ESTs	4.84
	448730	AB032983	Hs.21894	KIAA1157 protein	4.84
60	458339	AW976853	Hs.172843	ESTs	4.83
	422083	NM_001141	Hs.111256	arachidonate 15-lipoxygenase, second typ	4.82
	420159	Al572490	Hs.99785	Homo sapiens cDNA: FLJ21245 fis, clone C	4.82 4.82
	424103	NM_001918	Hs.139410	dihydrolipoamide branched chain transacy	4.82
65	449535 422048	W15267 NM 012445	Hs.23672 Hs.288126	low density lipoprotein receptor-related spondin 2, extracellular matrix protein	4.82
U.S	422048	AF154335	Hs.79691	LIM domain protein	4.82
	419972	AL041465	Hs.294038	golgin-67	4.81
	420235	AA256756	Hs.31178	ESTs	4.81
	423412	AF109300	Hs.147924	prostate cancer associated protein 6	4.80

	429598	AA811257	Hs.269710	ESTs	4.80
	457114	AJ821625	Hs.191602	ESTs	4.80
	421828 424602	AW891965 AK002055	Hs.289109 Hs.301129	histone deacetylase 3 hypothetical protein FLJ11193	4.79 4.78
5	428364	AA426565	Hs.160541	ESTs, Moderately similar to ALU1_HUMAN A	4.78
-	452335	AW188944	Hs.61272	ESTs	4.78
	410765	Al694972	Hs.66180	nucleosome assembly protein 1-like 2	4.77
	421040	AA715026	Hs.135280	ESTs	4.76
10	421518 452560	AI056392 BE077084	Hs.206819	ESTs ESTs	4.76 4.76
10	409752	AW963990		gb:EST376063 MAGE resequences, MAGH Homo	4.75
	439703	AF086538	Hs.198245	ESTs	4.75
	418836	Al655499	Hs.161712	ESTs	4.74
15	450642	R39773	Hs.7130	copine IV	4.74
13	419879 411440	Z17805 AW749402	Hs.93564	Homer, neuronal immediate early gene, 2 gb:QV4-BT0383-281299-061-c06 BT0383 Homo	4.74 4.74
	450649	NM 001429	Hs.297722	E1A binding protein p300	4.74
	408738	NM_014785	Hs.47313	KIAA0258 gene product	4.73
••	435020	AW505076	Hs.301855	DiGeorge syndrome critical region gene 8	4.72
20	411624	BE145964		KIAA0594 protein	4.72
	439360 440491	AA448488 R35252	Hs.55346 Hs.24944	ribosomal protein L44 ESTs, Weakly similar to 2109260A B cell	4.72 4.72
	442611	BE077155	Hs.177537	hypothetical protein DKFZp761B1514	4.72
	443555	N71710	Hs.21398	ESTs, Moderately similar to A Chain A, H	4.72
25	453900	BE300741	Hs.288416	hypothetical protein FLJ13340	4.72
	457528	AW973791	Hs.292784	ESTs	4.72
	416795	Al497778	Hs.168053	HBV pX associated protein-8	4.71 4.71
	407302 404721	R74206	Hs.268755	ESTs, Weakly similar to 178885 serine/th	4.70
30	426261	AW242243	Hs.168670	peroxisomal famesylated protein	4.70
	431924	AK000850	Hs.272203	Homo sapiens cDNA FLJ20843 fis, clone AD	4.70
	435256	AF193766	Hs.13872	cytokine-like protein C17	4.70
	438295	Al394151	Hs.37932	ESTs	4.70 4.70
35	442655 415788	AW027457 AW628686	Hs.30323 Hs.78851	ESTs, Weakly similar to B34087 hypotheti KIAA0217 protein	4.69
55	442760	BE075297	Hs.10067	ESTs, Weakly similar to A43932 mucin 2 p	4.69
	432432	AA541323	Hs.115831	ESTs	4.68
	454398	AA463437	Hs.11556	Homo sapiens cDNA FLJ12566 fis, clone NT	4.68
40	452741	BE392914	Hs.30503	Homo sapiens cDNA FLJ11344 fis, clone PL	4.67
40	424853 419706	BE549737 C04649	Hs.132967 Hs.77899	Human EST clone 122887 mariner transposo tropomyosin 1 (alpha)	4.67 4.66
	412088	Al689496	Hs.108932	ESTs	4.65
	416276	U41060	Hs.79136	LIV-1 protein, estrogen regulated	4.64
	429281	AAB30856	Hs.29808	Homo saplens cDNA: FLJ21122 fis, clone C	4.64
45	448207	AI475490	Hs.170577	ESTs	4.64
	408374 447162	AW025430 BE328091	Hs.155591 Hs.157396	forkhead box F1 ESTs, Weakly similar to A46010 X-linked	4.64 4.64
	451900	AB023199	Hs.27207	KIAA0982 protein	4.63
	421437	AW821252	Hs.104336	hypothetical prolein	4.63
50	418624	AI734080	Hs.104211	ESTs	4.63
	426172	AA371307	Hs.125056	ESTs	4.62
	439631 452994	AW136488 AW962597	Hs.25545 Hs.31305	ESTs KIAA1547 protein	4.61 4.61
	457726	Al217477	Hs.194591	ESTs	4.60
55	434629	AA789081	Hs.4029	glioma-amplified sequence-41	4.60
	403764			• •	4.58
	410659	Al080175	Hs.68826	ESTs	4.58
	432383 451246	AK000144 AW189232	Hs.274449 Hs.39140	Homo saplens cDNA FLJ20137 fis, clone CO cutaneous T-cell lymphoma tumor antigen	4.58 4.58
60	433234	AB040928	Hs.65366	KIAA1495 protein	4.57
-	424983	Al742434	Hs.169911	ESTs	4.56
	437812	AI582291	Hs.16846	ESTs, Weakly similar to O4HUD1 debrisoqu	4.56
	433447	AI082883	Hs.167593	hypothetical protein FLJ13409; KIAA1711	4.55
65	434715	BE005346	Hs.116410	ESTs	4.55
U.S	447673 408897	Al823987 N50204	Hs.182285 Hs.283709	ESTs lipopolysaccharide specific response-7 p	4.54 4.54
	438645	AW023424	Hs.156520	ESTs	4.54
	421247	BE391727	Hs.102910	general transcription factor IIH, polype	4.53
	450377	AB033091	Hs.24936	KIAA1265 protein	4.53
				152	
				152	

	433644 408321	AW405882	Hs.256112 Hs.44205	gb:hb75d03.x1 NCI_CGAP_Ut2 Homo saplens	4,53 4,53
	439225	AN900882 AA192869	Hs.45032	cortistatin ESTs	4.52
	440348	AW015802	Hs.47023	ESTs	4.52
5	446351	AW444551	Hs.258532	x 001 protein	4.52
-	451212	AW902672	Hs.287334	ESTs	4.52
	430294	A1538226	Hs.135184	guanine nucleotide binding protein 4	4.52
	435005	U80743	Hs.4316	trinucleotide repeat containing 12	4.52
10	448072	AI459306	Hs.24908	ESTs	4.50
10	403721 451018	AUGUSTON	Hs.247324	mitochondrial ribosomal protein S14	4.50 4.50
	451018	AW965599 AK001465	Hs.31575	SEC63, endoplasmic reticulum translocon	4.49
	417412	X16896	Hs.82112	interleukin 1 receptor, type I	4.48
	439735	AI635386	Hs.142846	hypothetical protein	4.48
15	435663	AI023707	Hs.134273	ESTs	4.48
	424036	AA770688	Hs.81946	H2A histone family, member L	4.48
	426388	AA748850	Hs.174877	bladder cancer overexpressed protein	4.48
	408622	AA056060	Hs.202577	Homo sapiens cDNA FLJ12166 fis, done MA	4.47 4.47
20	444269 430187	Al590346 Al799909	Hs.146220 Hs.158989	ESTs ESTs	4.46
20	427761	AA412205	Hs.140996	ESTs	4.46
	430261	AA305127	Hs.237225	hypothetical protein HT023	4.46
	444169	AV648170	Hs.58756	ESTs	4.44
	430598	AK001764	Hs.247112	hypothetical protein FLJ10902	4.44
25	412903	BE007967	Hs.155795	ESTs	4.44
	417048	AI068775	Hs.55498	geranylgeranyl diphosphate synthase 1	4.44
	442710	AI015631	Hs.23210	ESTs	4.44
	457413 400303	AA743462 AA242758	Hs.165337 Hs.79136	ESTs LIV-1 protein, estrogen regulated	4.42
30	443268	AJ800271	Hs.129445	hypothetical protein FLJ12496	4.42
30	438209	AL120659	Hs.6111	aryl-hydrocarbon receptor nuclear transl	4.42
	431724	AA514535	Hs.283704	ESTs	4.41
	412280	AW205116	Hs.272814	hypothetical protein DKFZp434E1723	4.40
~ ~	440801	AA906386	Hs.190535	ESTs	4.40
35	452959	AJ933416	Hs.189674	ESTs	4.40
	453861 417421	Al026838 AL138201	Hs.30120 Hs.82120	ESTs, Weekly similar to NUCL_HUMAN NUCLE nuclear receptor subfamily 4, group A, m	4.40
	417421	AC002551	Hs.331	general transcription factor IIIC, polyp	4.38
	433641	AF080229	113,001	ab:Human endogenous retrovirus K clone 1	4.38
40	447078	AW885727	Hs.301570	ESTs .	4.38
	424242	AA337476		hypothetical protein MGC13102	4.37
	408170	AW204516	Hs.31835	ESTs	4.38
	448757	Al366784	Hs.48820	TATA box binding protein (TBP)-associate	4.36 4.36
45	420021	AA252848	Hs.293557 Hs.253302	ESTs ESTs	4.36
43	449694 453867	A1659790 A1929383	Hs.108196	hypothetical protein DKFZp434N185	4.36
	458712	Al347502	Hs.173066	hypothetical protein FLJ20761	4.36
	417251	AW015242	Hs.99488	ESTs, Weakly similar to YK54_YEAST HYPOT	4.35
	434423	NM_006769	Hs.3844	LIM domain only 4	4.35
50	423427	AL137612	Hs.285848	KIAA1454 protein	4.34
	415715	F30364		ESTs	4.33 4.32
	404561	4.4700500	11- 400047	N-myrisloyitransferase 2	4.32
	422969 423685	AA782538 BE350494	Hs.122647 Hs.49753	uveal autoantigen with colled coll domai	4.32
55	443977	AL120986	Hs.150627	ESTs, Weakly similar to 138022 hypotheti	4.32
55	425071	NM_013989	Hs.154424	delodinase, iodothyronine, type II	4.32
	431583	AL042613	Hs.262476	S-adenosylmethlonine decarboxylase 1	4.31
	411379	AI816344	Hs.12554	ESTs, Weakly similar to NPL4_HUMAN NUCLE	4,30
	421476	AW953805	Hs.21887	ESTs	4.30
60	425178	H16097	Hs.161027	ESTs ESTs	4.30 4.30
	439262	AA832333	Hs.124399	hypothetical protein FLJ10879	4.30
	442818 421977	AK001741 W94197	Hs.8739 Hs.110165	ribosomal protein L26 homolog	4.29
	437114	AA836641	Hs.163085	ESTs	4.28
65	420195	N44348	Hs.300794	Homo sapiens cDNA FLJ11177 fis, clone PL	4.28
	418330	BE409405	Hs.94722	ESTs	4.27
	419750	AL079741	Hs.183114	Homo sapiens cDNA FLJ14236 fis, clone NT	4.28
	437065	AL036450	Hs.103238	ESTs gb:RC3-HT0585-160300-022-b09 HT0585 Homo	4.26 4.24
	455276	BE176479		ge.1105-1110000-100000-022-009 1110080 NOIRO	4.24

	416292	AA179233	Hs.42390	nasopheryngeal carcinoma susceptibility	4.24
	423740	Y07701	Hs.132243	aminopeptidase puromycin sensitive	4.24
	442023	AJ187878	Hs.144549	ESTs	4.24
	426764	AA732524	Hs.151464	ESTs, Weakly similar to ALUC_HUMAN !!!!	4.23
5	454058	AJ273419	Hs.135146	hypothetical protein FLJ 13984	4.23
	456511	AA282330	Hs.145668	ESTs	4.22
	448330	AL038449	Hs.207163	ESTs	4.22
	424701	NM_005923	Hs.151988	mitogen-activated protein kinase kinase	4.21
10	432621	AJ298501	Hs.12807	ESTs, Weakly similar to T46428 hypotheti	4.20
10	445707 419910	Al248720 AA662913	Hs.114390 Hs.190173	ESTs ESTs, Weakly similar to A46010 X-linked	4.20 4.20
	424085	NM 002914	Hs.139226	replication fector C (ectivetor 1) 2 (40	4.20
	440749	W22335	Hs.7392	hypothetical protein MGC3199	4.20
	442787	W93048	Hs.227203	hypothetical protein MGC2747	4.20
15	443414	R54594	Hs.25209	ESTs	4.20
	443558	AA256789	Hs.94949	methylmalonyl-CoA epimerase	4.20
	444170	AW613879	Hs.102408	ESTs	4.20
	446751	AA766998	Hs.85874	Human DNA sequence from clone RP11-16L21	4.20
20	421041	N36914	Hs.14691	ESTs, Mcderately similar to I38022 hypot	4.19
20	447476	BE293466	Hs.20880	ESTs, Weakly similar to 139022 hypotheti	4.19
	448543 410294	AW897741	Hs.21380 Hs.288891	Homo sapiens mRNA; cDNA DKFZp586P1124 (f	4.18
	433607	AB014515 AA602004	Hs.23260	KIAA0615 gene product ESTs	4.18 4.18
	435552	A1688636	Hs.193480	ESTs, Moderately similar to ALU6 HUMAN A	4.18
25	447124	AW976438	Hs.17428	RBP1-like protein	4.18
	453308	AW959731	Hs.32538	ESTs	4.17
	439328	W07411	Hs.118212	ESTs, Moderately similar to ALU3_HUMAN A	4.16
	430473	AW130690	Hs.299842	ESTs	4.16
	437257	AI283085	Hs.290931	ESTs, Weakly similar to YFJ7_YEAST HYPOT	4.16
30	438018	AK001160	Hs.5999	hypothetical protein FLJ10298	4.16
	443857	Al089292	Hs.287621	hypothetical protein FLJ14069	4.15
	446711 419103	AF169692 Z40229	Hs.12450	protocadherin 9	4.15 4.14
	405403	240229	Hs.96423	hypothetical protein FLJ23033	4.14
35	407378	AA299264		ESTs, Moderately similar to 138022 hypot	4.14
	408996	AW298602	Hs.197687	ESTs	4.14
	418727	AA227609	Hs.94834	ESTs	4.14
	434400	Al478211	Hs.186896	Homo sapiens cDNA FLJ11417 fis, clone HE	4.14
	438578	AA811244	Hs.164168	ESTs	4.14
40	450459	Al697193	Hs.299254	Homo sapiens cDNA: FLJ23597 fis, clone f.	4.14
	429887	AW366286	Hs.145696	splicing factor (CC1.3)	4.13
	448148 450316	NM_016578	Hs.20509 Hs.17850	HBV pX associated protein-8	4.13
	417531	W84446 NM 003157	Hs.1087	hypothetical protein MGC4543 serine/threonine kinase 2	4.12 4.12
45	431592	R69016	Hs.293871	hypothetical protein MGC10895s	4.12
	432463	AA548518	Hs.186733	ESTs	4.12
	433613	AA836126	Hs.5669	ESTs	4.12
	434739	AA804487	Hs.144130	ESTs	4.12
<b>~</b> 0	438259	AW205969	Hs.131808	ESTs	4.12
50	425810	AI923627	Hs.31903	ESTs	4.10
	432672	AW973775	Hs.130760	myosin phosphatase, target subunit 2	4.10
	433345	Al681545	Hs.152982 Hs.288031	hypothetical protein FLJ13117	4.10
	432712 453020	AB016247 AL162039	Hs.31422	sterol-C5-desaturase (fungal ERG3, delta Homo sapiens mRNA; cDNA DKFZp434M229 (fr	4.09 4.09
55	412045	AA099802	Hs.4299	transmembrane, prostate endrogen induced	4.09
-	435114	AA775483	Hs.288936	mitochondrial ribosomal protein L9	4.08
	443204	AW205878	Hs.29643	Homo sapiens cDNA FLJ13103 fis, clone NT	4.08
	445459	Al478629	Hs.158465	likely ortholog of mouse putative IKK re	4.08
	438938	H46212	Hs.137221	ESTs	4.07
60	454119	BE549773	Hs.40510	uncoupling protein 4	4.06
	411000	N40449	Hs.201619	ESTs, Weakly similar to S38383 SEB4B pro	4.06
	418926	AA232658	Hs.87070	UDP-glucose:glycoprotein glucosyltransfe	4.06
	424432 449673	AB037821 AA002064	Hs.146858 Hs.18920	protocadherin 10 ESTs	4.06 4.06
65	449073	AI620463	Hs.99197	hypothetical protein MGC13102	4.06
00	422174	AL049325	Hs.112493	Homo sapiens mRNA; cDNA DKFZp564D036 (fr	4.05
	455497	AA112573	Hs.285691	Homo sapiens prostein mRNA, complete cds	4.05
	415138	C18356	Hs.78045	tissue fector pethway inhibitor 2	4.04
	402791				4.04

	426792	AL044854	Hs.172329	KIAA0576 protein	4.04
	438660	U95740	Hs.6349	Homo sapiens, clone IMAGE:3010666, mRNA,	4.0
	442768	AL048534	Hs.48458	ESTs, Weakly similar to ALU8_HUMAN ALU S	4.0
-	447568	AF155655	Hs.18885	CGI-116 protein	4.0
5	428342	Al739168	Hs.131798	Homo saplens cDNA FLJ13458 fis, clone PL	4.0
	453439	Al572438	Hs.32976	guanine nucleotide binding protein 4	4.02
	453857	AL080235	Hs.35861	DKFZP686E1621 protein	4.02
	428249	AA130914	Hs.183291	zinc finger protein 268	4.02
10	432015	AL157504	Hs.159115	Homo sapiens mRNA; cDNA DKFZp58600724 (f	4.02
10	445495	BE622641	Hs.38489	ESTs, Weekly similar to 138022 hypotheti	4.02
	451746 452211	M88178 Al985513	Hs.233420	ESTs ESTs	4.02
	452211	AA284040	Hs.219441		4.02
	456038	AA203285	Hs.294141	ESTs, Highly similar to CA5B_HUMAN CARBO ESTs, Weakly similar to alternatively so	4.02
15	452449	AW068658	Hs.20943	ESTs veany similar to alternatively sp	4.02
13	407204	R41933	Hs.140237	ESTs. Weakly similar to ALU1 HUMAN ALU S	4.01
	428046	AW812795	Hs.155381	ESTs, Moderately similar to 138022 hypot	4.01
	438520	AA706319	Hs.98416	ESTs	4.01
	443292	AK000213	Hs.9196	hypothetical protein	4.01
20	432715	AA247152	Hs.200483	ESTs, Weakly similar to KIAA1074 protein	4.00
	403797	704.11104		and to the state of the state o	4.00
	418347	AA216419	Hs.289295	gb:nc16e03.s1 NCI_CGAP_Pr1 Homo sapiens	4.00
	419459	AW291128	Hs.278422	DKFZP586G1122 protein	4.00
	420911	U77413	Hs.100293	O-linked N-acetylglucosamine (GlcNAc) tr	4.00
25	425176	AW015844	Hs.301430	TEA domain family member 1 (SV40 transcr	4.00
	447505	AL049266	Hs.18724	Homo saplens mRNA; cDNA DKFZp564F093 (fr	4.00
	453773	AL133761		gb:DKFZp761C1413_r1 761 (synonym: harny2)	4.00
	434384	AA631910	Hs.162849	ESTs	3,99
	422471	AA311027	Hs.271894	ESTs, Weakly similar to (38022 hypotheti	3.99
30	427336	AW836261	Hs.177488	ESTs	3.98
	433394	Al907753	Hs.93810	cerebral cavernous malformations 1	3.98
	441269	AW015206	Hs.178784	ESTs	3.97
	419629	AB020695	Hs.91662	KIAA0888 protein	3.96
	435008	AF150262	Hs.182898	ESTs	3.96
35	456649	R74441	Hs.117178	poly(A)-binding protein, nuclear 1	3.96
	418723	AA504428	Hs.10487	Homo saplens, clone IMAGE:3954132, mRNA,	3.96
	428738	NM_000380	Hs.192803	xeroderma pigmentosum, complementation g	3.95
	430456	AA314998	Hs.241503	hypothetical protein	3.95
40	422017	NM_003877	Hs.110776	STAT induced STAT Inhibitor-2	3.95
40	409980	BE261944	Hs.153028	hexokinase 1	3.95
	455309	AW894017		gb:RC4-NN0027-150400-012-g04 NN0027 Homo	3.95
	450295	Al766732	Hs.201194	ESTs	3.94
	456660	AA909249	Hs.112282	solute carrier family 30 (zinc transport	3.94
45	410908 447145	AA121686 AA761073	Hs.10592 Hs.192943	ESTs	3.94
43	449318	AW236021		TRAF family member-associated NFKB activ Homo saplens, Similar to RIKEN cDNA 5730	3.94
	449839	W57990	Hs.108788 Hs.60059		3.94
	411887	AW182924	Hs.128790	Homo sapiens cDNA FLJ11478 fis, clone HE ESTs	3.93
	437531	AW182924 AI400752	Hs.112259	T cell receptor gamma locus	3.93
50	452238	F01811	Hs.187931	ESTs	3.93
20	410486	AW235094	Hs.193424	zinc finger protein	3.92
	424882	Al379461	Hs.153636	far upstream element (FUSE) binding prot	3.92
	426269	H15302	Hs.168950	Homo sapiens mRNA; cDNA DKFZp566A1046 (f	3.92
	427043	AA397679	Hs.298460	ESTs	3.92
55	440404	Al015881	Hs.125616	mitochandrial ribosomal protein S5	3.82
23	452762	AW501435	Hs.171409	v-akt murine thymoma viral oncogene homo	3.92
	453058	AW612293	Hs.288884	Homo sapiens cDNA FLJ11750 fls, clone HE	3.92
	423583	AL122055	Hs.129836	KIAA1028 protein	3.92
	408001	AA046458	Hs.95296	ESTs	3.92
60	419197	N48921	Hs.27441	KIAA1615 protein	3.91
	428635	Al355647	Hs.189999	purinergic receptor (family A group 5)	3.91
	401747			A	3.91
	410011	AB020641	Hs.57856	PFTAIRE protein kinase 1	3.91
	432205	AI806583	Hs.125291	ESTs	3.9
65	447857	AA061218	Hs.58608	Homo sapiens cDNA FLJ14206 fis, done NT	3.9
	448494	AA463276	Hs.288906	WW Domain-Containing Gene	3.91
	409928	AL137163	Hs.57549	hypothetical protein dJ473B4	3.90
		AL137163 BE336654	Hs.57549 Hs.70937	hypothetical protein dJ473B4 H3 histone femily, member A	3.90

	425707	AF115402	Hs.11713	E74-like factor 5 (ets domain transcript	3.90
	431325	AW026751	Hs.5794	ESTs, Weakly similar to 2109260A B cell	3.89
	451806	NM_003729	Hs.27076	RNA 3-terminal phosphate cyclase	3.89
-	401045				3.89
5	433023	AW864793	Hs.34161	thrombospondin 1	3.89
	452160	BE378541	Hs.279815	cysteine sulfinic acid decarboxylase-rel	3.89
	437372	AA323968	Hs.283631 Hs.81086	hypothetical protein DKFZp547G183	3.66
	417067 410467	AJ001417 AF102546	Hs.63931	solute carrier family 22 (extraneuronal dachshund (Drosophila) homolog	3.88
10	422660	AV102046 AW297582	Hs.237062	hypothetical protein FLJ22548 similar to	3.68
10	431930	AB035301	Hs.272211	cadherin 7, type 2	3.68
	453047	AW023798	Hs.286025	ESTs	3.88
	433891	AA613792	11020000	gb:no97h03.s1 NCI_CGAP_Pr2 Homo sapiens	3.88
	401785			•	3.88
15	431088	AA491824	Hs.196881	ESTs	3.88
	451952	AL120173	Hs.301663	ESTs	3.87
	422089	AA523172	Hs.103135	ESTs, Weakly similar to SFR4_HUMAN SPLIC	3.87
	452277	AL049013	Hs.28783	KIAA1223 protein	3.67
20	438279	AA305166	Hs.165165	HIV-1 rev binding protein 2	3.88
20	458229	Al929802	Hs.177	phosphatidylinositol glycan, class H	3.88
	406414			1	3.85
	417193	Al922189 AA723564	Hs.268390 Hs.191343	hypothetical protein FLJ22795 ESTs	3.85
	413174 433332	AA723004 Al367347	Hs.127809	Homo sapiens clone TCCCTA00151 mRNA sequ	3.85
25	411089	AA456454	Hs.118637	cell division cycle 2-like 1 (PITSLRE pr	3.85
23	412494	AL133900	Hs.792	ADP-ribosylation factor domain protein 1	3.84
	413530	AA130158	Hs.19977	ESTs, Moderately similar to ALU8_HUMAN A	3.84
	459592	AL037421	Hs.208746	ESTs, Moderately similar to pot. ORF I	3.84
	418329	AW247430	Hs.84152	cystathionine-beta-synthase	3.83
30	451468	AW503398	Hs.210047	ESTs, Moderately similar to I38022 hypot	3.83
	434804	AA649530		gb:ns44f05.s1 NCI_CGAP_Alv1 Homo sapiens	3.83
	401819				3.62
	424179	F30712		Homo sapiens, clone IMAGE:4285740, mRNA	3.82
25	424850	AA151057	Hs.153498	chromosome 18 open reading frame 1	3.82 3.82
35	426472	BE246138	Hs.30853	ESTS	3.62
	426625 427585	T78300 D31152	Hs.171409 Hs.179729	serologically defined colon cancer antig collagen, type X, alpha 1 (Schmid metaph	3.62
	427756	AI376540	Hs.15574	ESTs	3.82
	444701	Al916512	Hs.198394	ESTs	3.82
40	423052	M28214	Hs.123072	RAB3B, member RAS oncogene family	3.82
	429259	AA420450	Hs.292911	ESTs, Highly similar to S60712 band-6-pr	3.82
	416111	AA033813	Hs.79018	chromatin assembly factor 1, subunit A (	3.82
	433586	T85301		gb:yd78d06.s1 Soares fetal liver spleen	3.81
	438527	Al969251	Hs.143237	RAB7, member RAS oncogene family-like 1	3.81
45	410297	AA148710	Hs.159441	lumican	3.81
	429898	AW117322	Hs.42366	ESTs	3.81
	409079	W87707	Hs.82065	Interleukin 6 signal transducer (gp130,	3.80
	419423 429643	D26488 AA455889	Hs.90315 Hs.187548	KIAA0007 protein FYVE-finger-containing Rab5 effector pro	3.80
50	431499	NM_001514	Hs.258561	general transcription factor IIB	3.80
50	445060	AA830811	Hs.88808	ESTs	3.80
	449419	R34910	Hs.119172	ESTs	3.80
	450584	AA040403	Hs.60371	ESTs ·	3.80
	426137	AL040683	Hs.167031	DKFZP566D133 protein	3.79
55	420185	AL044058	Hs.158047	ESTs	3.79
	410076	T05387	Hs.7991	ESTs	3.78
	444078	BE246919	Hs.10290	U5 snRNP-specific 40 kDa protein (hPrp8-	3.78
	417318	AW953937	Hs.12891	ESTs	3.78
<b>c</b> 0	414664	AA587775	Hs.66295	multi-PDZ-domain-containing protein	3.78
60	410275	U85658	Hs.61796	transcription factor AP-2 gamma (activat	3.77 3.77
	410503 434170	AW975746 AA826509	Hs.188662 Hs.122329	KIAA1702 protein ESTs	3.77
	434170 421838	AW881089	Hs.122329 Hs.108806	Homo saplens mRNA; cDNA DKFZp566M0947 (f	3.77
	425268	Al807883	Hs.156932	Homo sapiens cDNA FLJ20653 fis, clone KA	3.76
65	431696	AA259068	Hs.267819	protein phosphatase 1, regulatory (inhib	3.76
0.0	411990	AW963624	Hs.31707	ESTs, Weakly similar to YEW4_YEAST HYPOT	3.76
	430291	AV660345	Hs.238126	CGI-49 protein	3.76
	448779	BE042877	Hs.177135	ESTs	3.76
	452632	AA456193	Hs.155606	progesterone membrane binding protein	3.75

3.75

	452598	AI831594	Hs.68647	ESTs, Weakly similar to ALU7_HUMAN ALU S	3.75
	439498	AA908731	Hs.58297	CLLL8 protein	3.75
	440258	Al741633	Hs.125350	ESTs	3.74
	456848	AL121087	Hs.296406	KIAA0685 gene product	3.74
5	415062	AA160000	Hs.137398	ESTs, Weakly similar to JC5238 galactosy	3.74
	420653	Al224532	Hs.88550	ESTs	3.74
	431637	AI879330	Hs.265960	hypothetical protein FLJ10563	3.74
	440411	N30256	Hs.156971	hypothetical protein DKFZp434G1415	3.74
	405917	1400200	110.100071	hypothetical protein DNI 2prova 1410	3.74
10	419440	AB020689	Hs.90419	KIAA0882 protein	3.74
10	451230	BE546208	Hs.26090	hypothetical protein FLJ20272	3.73
					3.73
	429597	NM_003816	Hs.2442	a disintegrin and metalloproteinase doma	
	430144	AI732722	Hs.187694	ERGL protein; ERGIC-53-like protein	3.72
10	438394	BE379623	Hs.27693	peptidylprolyl isomerase (cyclophilin)-l	3.72
15	440527	AV657117	Hs.184164	ESTs, Moderately similar to S65657 alpha	3.72
	449433	AI672096	Hs.9012	ESTs, Weakly similar to S26650 DNA-bindi	3.72
	456228	BE503227	Hs.134759	ESTs	3.72
	448663	BE614599	Hs.106823	hypothetical protein MGC14797	3.72
	415075	L27479	Hs.77889	Friedreich ataxia region gene X123	3.72
20	433544	Al793211	Hs.165372	ESTs, Moderately similar to ALU1_HUMAN A	3.71
	418293	Al224483	Hs.16063	hypothetical protein FLJ21877	3.71
	449897	AW819642	Hs.24135	transmembrane protein vezatin; hypotheti	3.71
	420297	Al628272	Hs.88323	ESTs, Weakly similar to ALU1_HUMAN ALU S	3.70
	423065	R96158	Hs.194606	Homo saplens, clone MGC:5406, mRNA, comp	3.70
25	429340	N35938	Hs.199429	Homo sapiens mRNA; cDNA DKFZp434M2216 (f	3.70
	437777	AA768098	Hs.189079	ESTs	3.70
	440351	AF030933	Hs.7179	RAD1 (S. pombe) homolog	3.70
	443603	BE502601	Hs.134289	ESTs, Weakly similar to KIAA1063 protein	3.70
	446965	BE242873	Hs.16677	WD repeat domain 15	3.70
30	412350	Al659306	Hs.73826	protein tyrosine phosphatase, non-recept	3.70
	433852	Al378329	Hs.126629	ESTs	3.70
	433142	AL120697	Hs.110640	ESTs	3.69
	419994	AA282881	Hs.190057	ESTs	3.69
	412628	AI972402	Hs.173902	hypothetical protein MGC2848	3.69
35	431416	AA532718	Hs.178804	ESTs	3.69
55	439444	Al277652	Hs.54578	ESTs, Weakly similar to I38022 hypotheti	3.68
	414709	AA704703	Hs.77031	Sp2 transcription factor	3.68
	447397	BE247676	Hs.18442	E-1 enzyme	3.68
	405718	DE247070	118, 10442	L-1 elizylile	3.68
40	425217	AU076696	Hs.155174	CDC5 (cell division cycle 5, S. pombe, h	3.68
40	442242		Hs.90424	Homo sapiens cDNA: FLJ23285 fis, done H	3.68
		AV647908			3.68
	424690	BE538356	Hs.151777	eukaryotic translation initiation factor	
	421734	Al318624	Hs.107444	Homo sapiens cDNA FLJ20562 fis, clone KA	3.67
45	427221	L15409	Hs.174007	von Hippel-Lindau syndrome	3.67
45	439864	Al720078	Hs.291997	ESTs, Weakly similar to A47582 B-cell gr	3.66
	402408				3.66
	426327	W03242	Hs.44898	Homo saplens clone TCCCTA00151 mRNA sequ	3.66
	427119	AW880562	Hs.114574	ESTs	3.66
~~	427356	AW023482	Hs.97849	ESTs	3.66
50	452946	X95425	Hs.31092	EphA5	3.66
	419078	M93119	Hs.89584	insulinoma-associated 1	3.66
	416235	AI064824	Hs.193385	ESTs	3.65
	427144	X95097	Hs.2126	vasoactive intestinal peptide receptor 2	3.65
	447500	AJ381900	Hs.159212	ESTs	3.65
55	453127	AI696671	Hs.294110	ESTs	3.65
	423396	Al382555	Hs.127950	bromodomain-containing 1	3.65
	419346	Al830417		polybramo 1	3.64
	441540	C01367	Hs.127128	ESTs	3.64
	446501	Al302616	Hs.150819	ESTs	3.64
60	459527	AW977556	Hs.291735	ESTs, Weakly similar to 178885 serine/th	3.63
	446320	AF126245	Hs.14791	acyl-Coenzyme A dehydrogenase family, me	3.63
	435706	W31254	Hs.7045	GL004 protein	3.63
	400110				3.62
	410313	R10305	Hs.185683	ESTs	3.62
65	414713	BE465243	Hs.12664	ESTs	3.62
	436279	AW900372	Hs.180793	ESTs, Weakly similar to \$65657 alpha-1C-	3.62
	439818	AL360137	Hs.19934	Homo sapiens mRNA full length insert cDN	3.62
	451797	AW663858	Hs.56120	small inducible cytokine subfamily E, me	3.62
	451294	AJ457338	Hs.29894	ESTs	3.62

452598 Al831594 Hs.68647 ESTs. Weakly similar to ALU7 HUMAN ALU S

	434194 404939	AF119847	Hs.283940	Homo sapiens PRO 1550 mRNA, partial cds
	408101	AW968504	Hs.123073	CDC2-related protein kinase 7
	435846	AA700870	Hs.14304	ESTs
5	432833	N51075	Hs.47191	ESTs
	427276	AA400269	Hs.49598	ESTs
	433495	AW373784	Hs.71	alpha-2-glycoprotein 1, zinc
	403137			
	404165			
- 10	409571	AA504249	Hs.187585	ESTs
	410561	BE540255	Hs.6994	Homo saplens cDNA: FLJ22044 fis, clone H
	412924	BE018422	Hs.75258	H2A histone family, member Y
	434228	Z42047	Hs.283978	Homo sapiens PRO2751 mRNA, complete cds
	436797	AA731491	Hs.178518	hypothetical protein MGC14879
15	437162	AW005505	Hs.5464	thyroid hormone receptor coactiveting pr
	437444	H46008	Hs.31518	ESTs
	404210			
	446157	BE270828	Hs.131740	Homo sapiens cDNA: FLJ22562 fis, clone H
20	437587	AJ591222	Hs.122421	Human DNA sequence from clone RP1-187J11
20	423147	AA987927	Hs.131740	Homo sapiens cDNA: FLJ22562 fis, clone H
	452226	AA024898	Hs.296002	ESTs
	443775	AF291664	Hs.204732	matrix metalloproteinase 26
	452501	AB037791	Hs.29716	hypothetical protein FLJ10980
25	428647	AA830050	Hs.124344	ESTs
23	422443	NM_014707	Hs.116753	histone deacetylase 78
	447966 420892	AA340605 AW975076	Hs.105887 Hs.172589	ESTs, Weakly similar to Homolog of rat Z
				nuclear phosphoprotein similar to S. cer
	420230 418428	AL034344 Y12490	Hs.298020	forkhead box C1
30			Hs.85092	thyroid hormone receptor interactor 11
30	428949 444929	AA442153 Ai685841	Hs.104744 Hs.161354	hypothetical protein DKFZp434J0617 ESTs
	433339	AF019226	Hs.8036	
	433339	R87622	Hs.26714	glioblastoma overexpressed
	433002	AF048730	Hs.279906	KIAA1831 protein cyclin T1
35	435425	H16263	Hs.31416	ESTs
33	415621	AI648602	Hs.131189	ESTs
	416974	AF010233	Hs.80667	RALBP1 associated Eps domain containing
	405793	APU 10233	ns.6000/	nALDF I associated the domain containing
	409770	AW499538		gb:Ul-HF-BR0p-aji-c-12-0-Ul.r1 NIH_MGC_5
40	425305	AA363025	Hs.155572	Human clone 23801 mRNA sequence
••	428939	AW238550	Hs.131914	ESTs
	438388	AA806349	Hs.44698	ESTs
	443703	AV646177	Hs.213021	ESTs
	457940	AL360159	Hs.30445	Homo saplens TRipartite motif protein ps
45	402444			
	409643	AW450868	Hs.257359	ESTs
	418250	U29926	Hs.83918	adenosine monophosphate deaminase (isofo
	432745	AI821926	Hs.269507	gb:nt78f05.x5 NCI_CGAP_Pr3 Homo saplens
	414222	AL135173	Hs.878	sorbitol dehydrogenase
50	430061	AB037817	Hs.230188	KIAA1396 protein
	421491	H99969	Hs.42736	ESTs
	422384	AA224077	Hs.42438	Sm protein F
	434565	T52172		ESTs
	438379	N23018	Hs.171391	C-terminal binding protein 2
55	439741	BE379646	Hs.6904	Homo sapiens mRNA full length insert cDN
	447311	R37010	Hs.33417	Homo sapiens cDNA: FLJ22806 fis, clone K
	447805	AW627932	Hs.19614	gemin4
	454265	H03556	Hs.300949	ESTs, Weakly similar to thyrold hormone
	418838	AW385224	Hs.35198	ectonucleotide pyrophosphatase/phosphodi
60	448804	AW512213	Hs.42500	ADP-ribosylation factor-like 5
	409617	BE003760	Hs.55209	Homo sapiens mRNA; cDNA DKFZp434K0514 (f
	434075	AW003416	Hs.160604	ESTs
	444190	AI878918	Hs.10526	cysteine and glycine-rich protein 2
	435017	AA338522	Hs.12854	angiotensin II, type I receptor-associat
65	423445	NM_014324	Hs.128749	alpha-methylacyl-CoA racemase
	420271	AI954365	Hs.42892	ESTs
	443684	Al681307	Hs.166674	ESTs
	444168 446074	AW379879 AA079799	Hs.29263	gb:RC1-HT0256-081199-011-f01 HT0256 Homo hypothetical protein FLJ11896

	452582 431542	AL137407 H63010	Hs.29911 Hs.5740	Homo sapiens mRNA; cDNA DKFZp434M232 (ir ESTs	3.48 3.48
	432697	AW975050	Hs.293692	ESTs, Weakly similar to ALU4_HUMAN ALU S	3.48
	435572	AW975339	Hs.239828	ESTs. Weakly similar to GAG2 HUMAN RETRO	3.47
5	407192	AA609200	110203020	gb:af12e02.s1 Soares testis NHT Homo sap	3.47
,	413435	X51405	Hs.75360	carboxypeptidase E	3,46
	447210	AF035269	Hs.17752	phosphatidviserine-specific phospholipas	3.46
	447210	AW796524	Hs.68644	Homo sapiens microsomal signal peptidase	3,46
	425312	ANV/90024 AA354940	Hs.145958	ESTs	3.46
10					
10	442007	AA301116	Hs.142838	nucleolar phosphoprotein Nopp34	3.46
	417455	AW007066	Hs.18949	ESTs, Weakly similar to CA2B_HUMAN COLLA	3.45
	426931	NM_003416	Hs.2076	zinc finger protein 7 (KOX 4, clone HF.1	3.45
	403739	W01556	Hs.238797	ESTs, Moderately similar to 138022 hypot	3.45
	436024	Al800041	Hs.190555	ESTs	3.45
15	403418	AW963897	Hs.44743	KJAA1435 protein	3.45
	409151	AA306105	Hs.50785	SEC22, vesicle trafficking protein (S. c.	3.44
	418626	AW299508	Hs.135230	ESTs	3.44
	420560	AW207748	Hs.59115	ESTs	3.44
	420686	Al950339	Hs.40782	ESTs	3.44
20	428870	AA436831	Hs.36049	ESTs	3.44
	436754	Al061288	Hs.133437	ESTs	3,44
	437960	A1669586	Hs.222194	ESTs	3,44
	452300	AW628045	Hs.28896	Homo sapiens mRNA full length insert cDN	3.44
	421887	AW161450	Hs.109201	CGI-86 protein	3.44
25	-12.100/	ATT 101400	10.100201	Odi oo piotoiii	3.44

TABLE 5A shows the accession numbers for those primekeys lacking a unigeneID in Tables 5, 6, and 7. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers for sequences comprising each cluster are listed in the "Accession" column.

	Pkey: CAT number: Accession:		Unique Eos probeset Identifier number Gene cluster number Genbank coesskon numbers
10	Accession.		Generality accession monitoris
	Pkey	CAT number	Accession
	407596	1003489_1	R86913 R86901 H25352 R01370 H43764 AW044451 W21298
	408432	1058667_1	AW195262 R27868 AW811262
15	409752	115301_1	AW963990 AA078196 AW749482 AA077488 BE151571 AA376917
	409770	1154048_1	AW499536 AW499553 AW502138 AW499537 AW502136 AW501743
	411440	124577_1	AW749402 AW749403 Z45743 R80376 AA093358
	411479	1247077_1	AW848047 AW848202 AW848631 AW848142 AW848702 AW848121 AW848632 AW848140 AW848571 AW848009 AW848067 AW848069 AW848905 AW848214
20	411624	1252166_1	BE145964 BE146286 AW854564
	412991	134248_1	AW949013 AA126111
	414269	143133_1	AA298489 AA137165
	415123	1523390_1	D60925 D60628 D80787
	415715	1548818_1	F30364 F36569 T15435
25	416288	1585983_1	H51299 H44619 H46391 R86024 H51892 T72744
	416289	1586037_1	W26333 R05358 H44682
	417730	1695795_1	Z44761 R25801 R11926 R35604
	418636	177402_1	AW749855 AA225995 AW750208 AW750206
	419346	184129_1	AI830417 AA236612
30	419536	185688_1	AA603305 AA244095 AA244183
	420111	190755_1	AA255652 AA260911 AW987920 AA262684
	422219	213547_1	AW978073 AW978072 AA807550 AA306567
	424179	236389_1	F30712 F35665 AW263888 AI904014 AI904018 AA336927 AA336502
	424242	237181_1	AA337476 AW966227 AA450376 AW960222 AA381051
35	428002	285602_1	AA418703 AA418711 BE071915 BE071920 BE071912
	429163	300543_1	AA884766 AW974271 AA592975 AA447312
	432189	342819_1	AA527941 AI810608 AI620190 AA635266
	432340	345248_1	AA534222 AA632632 T81234
••	432363	345469_1	AA534489 AW970240 AW970323
40	432966	356839_1	AA850114 AW974148 AA572946
	433586	370470_1	T85301 AW517087 AA601054 BE073959
	433641	37186_1	AF080229 AF080231 AF080230 AF080232 AF080233 AF080234 BE550533 Al636743 AW814951 BE467547
			Al680833 Al633818 N29988 U67592 U67593 U67590 U67591 S46404 U67567 AA463992 AW206802 Al970376
45			AI583718 AI672574 N25695 AW665466 AI818326 AA126128 AI480345 AW013827 AA246638 AI214968 AA2n4735 AA207155 AA206262 AA204833 AW003247 AW496808 AI080480 AI691703 AI651023 AI967418
ŧ5			AA204735 AA207155 AA203262 AA204833 AW003247 AW498806 Al080480 Al651703 Al651023 Al667416 AW818140 AA502500 Al206199 Al671282 Al352545 BE501030 Al652535 BE465762 AA206331 AW451866
			AW818140 AA502500 AI206199 AI671282 AI352545 BE501030 AI652535 BE465762 AA206331 AW451865  AA471088 AA206342 AA204834 AA206100 AW021661 AA332922 N66048 AA703396 H92278 AW139734
			H92683 U87589 U87595 H69001 U87594 BE466420 A/624817 BE466611 A/206344 AA574397 AA348354
50			AI493192
ou	433687	373061_1	AA743991 AA604852 AW272737
	433891	376239_1	AA613792 AW182929 T05304 AW858385
	434415 434565	385931_1	BE177494 AW276909 AA632849 T52172 AF147324 T52248
	434565	38898_1	AA649590 AA659316 H64973
55	434804 437113	393481_1	AA744693 AW750059
JJ		433234_1	AW379879 AH26285 H12014
	444168 448212	593829_1 755099_1	AW3/96/9 AH26285 F12014 AI475858 AW969013
	448212 448310	755099_1 757918_1	AI4/5858 AW959013 AI480316 AW847535
	448310 451746		M86178 Al813822 D56993
	431740	883303_1	M00110 VI010075 D00000

	452560	922216_1	BE077084 AW139963 AW863127 AW808209 AW806204 AW606205 AW806206 AW806211 AW806212 AW806207 AW806208 AW806210 AB07497
	452712 453773	928309_1 980699_1	AW838516 AW839560 BE144343 Al914520 AW888910 BE184854 BE184784 At 133761 At 133767
5	455276	1272541_1	BE176479 BE176676 BE176357 BE176550 AW886079 BE176676 BE176615 BE176655 BE176489 BE176610 BE176362
	455309	1278153_1	AW894017 AW893956 AW894032

TABLE 5B shows the genomic positioning for those primekeys lacking unigene ID's and accession numbers in Tables 5, 6, and 7. For each predicted exon, we have listed the

Unique number corresponding to an Eos probeset Sequence source. The 7 digit numbers in this column are Genbank Identifier (GI) numbers. "Dunham I. et al." refers to the

publication entitled "The DNA sequence of human chromosome 22." Dunham I. et al., Nature (1999) 402:489-495.

genomic sequence source used for prediction. Nucleotide locations of each predicted exon 5 are also listed.

10

	Strand: Nt_position:		Indicates DNA strand from which exons were predicted. Indicates nucleoticle positions of predicted exons.					
15								
	Pkey	Ref	Strand	Nt_position				
	401045	8117819	Plus	90044-90184,91111-91345				
20	401424	8176894	Plus	24223-24428				
	401451	6634068	Minus	119926-121272				
	401714	6715702	Plus	96484-98681				
	401747	9769672	Minus	118596-118816,119119-119244,119609-119761,120422-120990,130161-130381,130468-130593,131097- 131258,131866-131932,132451-132575,133580-134011				
25	401785	7249190	Minus	165776-165996,166189-166314,166408-166569,167112-167268,167387-167469,168634-168942				
	401819	7467933	Minus	28217-28496				
	402408	9796239	Minus	110326-110491				
	402444	9796614	Plus	28391-28517				
	402791	6137008	Minus	51036-51207				
30	403047	3540153	Minus	59793-59968				
	403137	9211494	Minus	92349-92572,92958-93084,93579-93712,93949-94072,94591-94748,95214-95337				
	403721	7528046	Minus	156647-157366				
	403764	7717105	Minus	118692-118853				
	403797	8099896	Minus	123065-125008				
35	404165	9926489	Minus	69025-69128				
	404210	5006246	Plus	169928-170121				
	404253	9367202	Minus	55675-56055				
	404561	9795980	Minus	69039-70100				
40	404571	7249169	Minus	112450-112648				
40	404721	9856648	Minus	173763-174294				

Minus

Minus Minus

Minus

100915-101087 175318-175476

37956-38097

89197-89453

39694-40031

49593-49850 106956-107121

106829-107213

113060-113266

37491-37870,40951-41031

404915 7341766

404939 6862697 Plus

405403 6850244

405685 4508129

405718 9795467 Plus 1405887

405793

405876 6758747 Plus

406414 9258407 Plus

406554 7711566 Plus

405917 7712162 Minus

45

## TABLE 6:286 GENES ENCODING EXTRACELLULAR OR CELL SURFACE PROTEINS UP-REGULATED IN PROSTATE CANCER COMPARED TO NORMAL ADULT TISSUES

Table 6 shows 286 genes up-regulated in prostate cancer compared to normal adult tissues 5 that are likely to be extracellular or cell-surface proteins. These were selected as for Table 5 and the predicted protein contained a structural domain that is indicative of extracellular localization (e.g. egf, 7tm domains).

	Pkey: ExAccn: UnigeneID: Unigene Title: R1:		Unique Eos probeset identifier number Exempter Accession number, Genbank accession number Unique neumber Unique gene tible Alabo of tumor to normal fissue				
10	Pkey	ExAcon	UnigeneID	Uningene Title	R1_		
	409361 409731	NM_005982	Hs.54416	sine oculis homeobox (Drosophila) homolo	48.28		
		AA125985	Hs.56145	thymosin, beta, identified in neuroblast	45.24		
15	400298	AA032279	Hs.61635	six transmembrane epithelial antigen of	43.48		
13	420154	Al093155	Hs.95420	JM27 protein	41.12		
	426747	AA535210	Hs.171995	kallikrein 3, (prostate specific entigen	31.80		
	400299	X07730	Hs.171995	kallikrein 3, (prostate specific entigen	24.91		
	425075	AA506324	Hs.1852	acid phosphatase, prostate	24.23		
20	424846	AU077324	Hs.1832	neuropeptide Y	23.57		
20	405685	Vennen			20.90		
	420757	X78592	Hs.99915	androgen receptor (dihydrotestosterone r	19.72		
	418994	AA296520	Hs.89546	selectin E (endothelial adhesion molecul	19.56		
	452792	AB037765	Hs.30652	KIAA1344 protein	17.39		
25	445472	AB006631	Hs.12784	Homo sapiens mRNA for KIAA0293 gene, par	17.00		
23	414565	AA502972	Hs.183390	hypothetical protein FLJ13590	16.82		
	431716	D89053	Hs.268012	fatty-acid-Coenzyme A ligase, long-chain	16.60		
	408430	S79876	Hs.44926	dipeptidylpeptidase IV (CD26, adenosine	16.28		
	408000	L11690	Hs.620	bullous pemphigoid antigen 1 (230/240kD)	15.54		
30	430226	BE245562	Hs.2551	adrenergic, beta-2-, receptor, surface	15.40		
<b>3</b> 0	444484	AK002126	Hs.11260	hypothetical protein FLJ11264	14.76		
	418601 448999	AA279490	Hs.86368	calmegin	14.56		
		AF179274	Hs.22791	transmembrane protein with EGF-like and	14.55		
	416182 420544	NM_004354 AA677577	Hs.79069	cyclin G2 Homo sapiens Chromosome 16 BAC clone CIT	12.94 12.79		
35	445413	AA151342	Hs.98732 Hs.12677		12.79		
33	453930	AA191342 AA419466	Hs.12677 Hs.36727	CGI-147 protein	12.04		
	440286	U29589	Hs.7138	hypothetical protein FLJ10903	12.04		
	452784	BE463857	Hs.151258	cholinergic receptor, muscarinic 3 hypothetical protein FLJ21062	11.88		
	452704	AF097994	Hs.301528		11.68		
40	448045	AJ297436	Hs.20166	L-kynurenine/alpha-aminoadipate aminotra prostate stem cell antigen	11.51		
40	449650	AF055575	Hs.23838	calcium channel, voltage-dependent, L tv	11.18		
	420381	D50640	Hs.337616	phosphodiesterase 3B, cGMP-inhibited	11.10		
	425665	AK001050	Hs.159066	hypothetical protein FLJ10188	11.08		
	425710	AF030880	Hs.159275	solute carrier family, member 4	11.08		
45	428728	NM 016625	Hs.191381	hypothetical protein	11.04		
43	407021	U52077	na. 19 100 1	gb:Human mariner1 transposase gene, comp	11.02		
	410733	D84284	Hs.66052	CD38 antigen (p45)	11.02		
	452340	NM 002202	Hs.505	ISL1 transcription factor, LIM/homeodoma	10.85		
	428819	AL135623	Hs.193914	KIAA0575 gens product	10.48		
50	421991	NM_014918	Hs.110488	KIAA0990 protein	10.04		
55	431217	NM_013427	Hs.250830	Rho GTPase activating protein 6	9.75		
	421470	R27496	Hs.1378	annexin A3	9.64		
	409262	AK000631	Hs.52256	hypothetical protein FLJ20624	9.45		
	435980	AF274571	Hs.129142	deoxyribonuclease II beta	9.24		
55	421246	AW582962	Hs.102897	CGI-47 protein	9.20		
55	410001	AB041036	Hs.57771	kallikrein 11	9.03		
	441791	AW372449	Hs.175982	hypothetical protein FLJ21159	9.02		

8.66

	404571				8.66
	456497	AW967956	Hs.123648	ESTs, Weakly similar to AF108460 1 ubinu	8.56
	419968	X04430	Hs.93913	interleukin 6 (interferon, beta 2)	8.36
	433172	AB037841	Hs.102652	hypothetical protein ASH1	8.30
5	422631	BE218919	Hs.118793	hypothetical protein FLJ10688	8.27
	427674	NM_003528	Hs.2178	H2B histone family, member Q	8.20
	404915				8.03
	452259	AA317439	Hs.28707	signal sequence receptor, gamma (translo	8.06
	452891	N75582	Hs.212875	ESTs, Weakly similar to DYH9_HUMAN CILIA	8.02
10	439731	A/953135	Hs.45140	hypothetical protein FLJ14084	7.98
	419839	U24577	Hs.93304	phospholipase A2, group VII (platelet-ac	7.68
	420120	AL049810	Hs.95243	transcription elongation factor A (SII)-	7.64
	424099	AF071202	Hs.139338	ATP-binding cassatte, sub-family C (CFTR	7.64
	448706	AW291095	Hs.21814	interleukin 20 receptor, alpha	7.52
15	410227	AB009284	Hs.61152	exostoses (multiple)-like 2	7.49
	425211	M18667	Hs.1867	progastricsin (pepsinogen C)	7.35
	441736	AW292779	Hs.169799	ESTs	7.28
	419991	AJ000098	Hs.94210	eyes absent (Drosophila) homolog 1	7.20
	425018	BE245277	Hs.154196	E4F transcription factor 1	7.20
20	424560	AA158727	Hs.150555	protein predicted by clone 23733	7.18
20	409110	AA191493	Hs.48778	niben protein	7.10
	421566	NM 000399	Hs.1395	early growth response 2 (Krox-20 (Drosop	7.04
	431725	X65724	Hs.2839	Norrie disease (pseudoglioma)	6.98
	425782	U66468	Hs.159525	cell growth regulatory with EF-hand doma	6.85
25	427408	AA583206	Hs.2156	RAR-related orphan receptor A	6.79
25	435604	AA625279	Hs.26892	uncharacterized bone marrow protein BM04	6.73
	415874	AF091622	Hs.78893	KIAA0244 protein	6.54
	401451	AI USTOLL	110.70000	TOP COLUMN PROCESS	6.52
	431778	AL080276	Hs.268562	regulator of G-protein signalling 17	6.51
30	409089	NM 014781	Hs.50421	KIAA0203 gene product	6.50
50	431992	NM_002742	Hs.2891	protein kinase C, mu	8.49
	401253	NM_002742	H8.2001	protein kinase o, ma	6.42
	404253	AF026692	Hs.105700	secreted frizzlad-related protein 4	6.41
	416806	NM 000288	Hs.79993	peroxisomal biogenesis factor 7	6.38
35		X63629	Hs.2877	cadherin 3, type 1, P-cadherin (placenta	6.30
55	431958 439366	AF100143	Hs.6540	fibroblast growth factor 13	8.30
	416836	D54745	Hs.80247	cholecystokinin	6.30
			Hs.192731	double-stranded RNA specific adenosine d	6.29
	433383 450728	AF034837 AW162923	Hs.25363	presenilin 2 (Alzheimer disease 4)	6.25
40	413384	NM 000401	Hs.75334	exostoses (multiple) 2	6.22
40	423349	AF010258	Hs.127428	homeo box A9	6.20
	424800	AL035588	Hs.153203	MyoD family inhibitor	6.18
	425451	AF242769	Hs.157461	mesenchymal stem cell protein DSC54	6.14
	447359	NM 012093	Hs.18268	adenviate kinese 5	6.00
45	410889	X91662	Hs.66744	twist (Drosophila) homolog (acrocephalos	5.97
43		NM 006042	Hs.48384	heparan sulfate (glucosamine) 3-O-sulfot	5.94
	408829 453911	AW503857	Hs.4007	Sarcolammal-associated protein	5.94
			Hs.48604	DKFZP434B168 protein	5.92
	408875	NM_015434 X82125	Hs.25040	zinc finger protein 239	5.90
50	450480 451684	AF216751	Hs.26813	CDA14	5.88
50	400301	X03835	Hs.1657	estrogen receptor 1	5.78
			Hs.934	glucosaminyi (N-acetyl) transferase 2, I	5.74
	415077	L41607 BE537037	Hs.273294	hypothetical protein FLJ20069	5.72
	418852	AB007891	Hs.16349	KIAA0431 protein	5.72
55	446867		Hs.61184	CGI-79 protein	5.70
33	410232	AW372451	Hs.119976	Human DNA sequence from clone RP1-20N2 o	5.70
	422762	AL031320	Hs.302689	hypothetical prolein	5.70
	450616	AL133067	Hs.46638		5.65
	408621	A1970672	Hs.6641	chromosome 11 open reading frame 8	5.64
60	439671	AW162840	Hs.59838	kinesin family member 5C hypothetical protein FLJ10808	5.60
00	410196	A1936442			5.60
	429170	NM_001394	Hs.2359	dual specificity phosphatase 4	5.60
	440738	AI004650	Hs.225674	WD repeat domain 9	5.59
	414342	AA742181	Hs.75912	KIAA0257 prolein	5.56
	422634	NM_016010	Hs.118821	CGI-62 protein	5.55
65	400288		11: 000000	050041-1-	5.51
	439569	AW602186	Hs.222399	CEGP1 protein	5.48
	452823	AB012124	Hs.30696	transcription factor-like 5 (basic helix	5.44
	431938	AA938471	Hs.54431 Hs.208341	specific granule protein (28 kDa); cyste	5.42
	427638	AA406411	HS208341	ESTs, Weakly similar to KIAA0989 protein	5.42

404571

	421264	AL039123	Hs.103042	microtubule-associated protein 1B	5.38
	421685	AF189723	Hs.106778	ATPase, Ca++ transporting, type 2C, memb	5.37
	421987	Al133161	Hs,286131	CGI-101 protein	5.36
-	422806	BE314767	Hs.1581	glutathione S-transferase theta 2	5.34
5	432281	AK001239	Hs.274263	hypothetical protein FLJ10377 Homo sapiens mRNA; cDNA DKFZp564O1763 (f	5.32 5.32
	451982 444042	F13036 NM 004915	Hs.27373 Hs.10237	ATP-binding cassette, sub-family G (WHIT	5.31
	444042 447752	M73700	Hs.10237	lactotransferrin	5.29
	451418	BE387790	Hs.26389	hypothetical protein FLJ20287	5.22
10	428593	AW207440	Hs.185973	degenerative spermatocyte (homolog Droso	5.21
10	447541	AK000288	Hs.18800	hypothetical protein FLJ20281	5.18
	459294	AW977286	Hs.17428	RBP1-like protein	5.16
	424692	AA429834	Hs.151791	KIAA0092 gene product	5.15
	416434	AW163045	Hs.79334	nuclear factor, interleukin 3 regulated	5.11
15	410268	AA316181	Hs.61635	six transmembrane epithelial antigen of	5.10
	417517	AF001176	Hs.82238	POP4 (processing of precursor, S. cerev	5.10
	453616	NM_003462	Hs.33846	dynein, axonemal, light intermediate pol	5.10
	427958	AA418000	Hs.96280	potassium intermediate/small conductance	5.09 5.03
20	407945	X69208	Hs.606	ATPase, Cu++ transporting, alpha polypep	5.05
20	418576 413328	AW968159 Y15723	Hs.289104 Hs.78295	Alu-binding protein with zinc finger dom quanylate cyclase 1, soluble, alpha 3	5.04
	432729	AK000292	Hs.278732	hypothetical protein FLJ20285	5.04
	432729	AF093419	Hs.169378	multiple PDZ domain protein	5.02
	429782	NM 005754	Hs.220689	Ras-GTPase-activating protein SH3-domain	5.02
25	436209	AW850417	Hs.254020	ESTs. Moderately similar to unnamed prot	5.02
	430599	NM 004855	Hs.247118	phosphatidylinositol glycan, class B	5.00
	451386	AB029006	Hs.26334	spastic paraplegia 4 (autosomal dominant	5.00
	457211	AW972565	Hs.32399	ESTs, Weakly similar to S51797 vasodilat	4.97
	425851	NM_001490	Hs.159642	glucosaminyl (N-acetyl) transferase 1, c	4.97
30	421689	N87820	Hs.106826	KIAA1696 protein	4.93
	416533	BE244053	Hs.79362	retinoblastoma-like 2 (p130)	4.92 4.91
	432653 403047	N62096	Hs.293185	ESTs, Weakly similar to JC7328 amino aci	4.91
	403047	AF003522	Hs.250500	delta (Drosophila)-like 1	4.90
35	427617	D42063	Hs.199179	RAN binding protein 2	4.88
55	428804	AK000713	Hs.193736	hypothetical protein FLJ20706	4.88
	449071	NM_005872	Hs.22960	breast carcinoma amplified sequence 2	4.86
	407596	R86913		gb:yq30f05.r1 Soares fetal liver spleen	4.84
	456516	BE172704	Hs.222746	KIAA1610 protein	4.84
40	458339	AW976853	Hs.172843	ESTs	4.83
	422083	NM_001141	Hs.111258	arachidonate 15-lipoxygenase, second typ	4.82
	449535	W15267	Hs.23672	low density lipoprotein receptor-related	4.82
	422048	NM_012445	Hs.268128	spondin 2, extracellular matrix protein	4.82 4.78
45	424602 410765	AK002055 Al694972	Hs.151046 Hs.66180	hypothetical protein FLJ11193 nucleosome assembly protein 1-like 2	4.77
43	410/65	Z17805	Hs.93564	Homer, neuronal immediate early gene, 2	4.74
	450649	NM_001429	Hs.25272	E1A binding protein p300	4.74
	411624	BE145964	Hs.103283	KIAA0594 protein	4.72
	404721	0211000			4.70
50	426261	AW242243	Hs.168670	peroxisomal famesylated protein	4.70
	416276	U41060	Hs.79136	LIV-1 protein, estrogen regulated	4.64
	403374	AW025430	Hs.155591	forkhead box F1	4.64
	451900	AB023199	Hs.27207	KIAA0982 protein	4.63
	421437	AW821252	Hs.104336	hypothetical protein	4.63
55	434629	AA789081	Hs.4029	glioma-ampilfied sequence-41	4.60 4.58
	403764 421247	BE391727	Hs.102910	general transcription factor IIH, polype	4.53
	403721	BE391727	us'insain	general transcription factor firs, potype	4.50
	453070	AK001465	Hs.31575	SEC63, endoplasmic reticulum translocon	4,49
60	417412	X16896	Hs.82112	interleukin 1 receptor, type I	4.48
50	439735	AI635386	Hs.142846	hypothetical protein	4.48
	430261	AA305127	Hs.237225	hypothetical protein HT023	4.46
	430598	AK001764	Hs.247112	hypothetical protein FLJ10902	4.44
	400303	AA242758	Hs.79136	LIV-1 protein, estrogen regulated	4.42
65	438209	AL120659	Hs,6111	aryl-hydrocarbon receptor nuclear transl	4.42
	417421	AL138201	Hs.82120	nuclear receptor subfamily 4, group A, m	4.40 4.38
	447270 434423	AC002551	Hs.331 Hs.3844	general transcription factor IIIC, polyp LIM domain only 4	4.35
	434423 404561	NM_006769	MS.3844	Eim domain only 4	4.32
	404001				

	422989	AA782538	Hs.122647	N-myristoyltranslerase 2	4.32
	423685	BE350494	Hs.49753	uveal autoanligen with colled coil domai	4.32
	425071	NM_013989	Hs.154424	delodinase, iodothyronine, type li	4.32
	431583	AL042613	Hs.262476	S-adenosylmethionine decarboxylase 1	4.31
5	442818	AK001741	Hs.8739	hypothetical protein FLJ10879	4.30
	423740	Y07701	Hs.293007	aminopeptidase puromycin sensitive	4.24
	424701	NM_005923	Hs.151988	mitogen-activated protein kinase kinase	4.21
	424085	NM_002914	Hs.139226	replication factor C (activator 1) 2 (40	4.20
	410294	AB014515	Hs.323712	KIAA0615 gene product	4.18
10	447124	AW976438	Hs.17428	RBP1-like protein	4.18
	438018	AK001160	Hs.5999	hypothetical protein FLJ10298	4.16
	443857	Al089292	Hs.287621	hypothetical protein FLJ14069	4.15
	446711	AF169692	Hs.12450	protocadherin 9	4.15
	405403				4.14
15	448148	NM_016578	Hs.20509	HBV pX associated protein-8	4.13
	417531	NM_003157	Hs.1087	serine/threonine kinase 2	4.12
	433345	Al681545	Hs.152982	hypothetical protein FLJ13117	4.10
	432712	AB016247	Hs.288031	sterol-C5-desaturase (fungal ERG3, delta	4.09
	435114	AA775483	Hs.288936	milochondrial ribosomal protein L9	4.08
20	445459	Al478629	Hs.158465	fixely ortholog of mouse putative IKK re	4.08
	402791				4.04
	438660	U95740	Hs.6349	Homo sapiens, done IMAGE:3010668, mRNA,	4.04
	447568	AF155655	Hs.18885	CGI-116 protein	4.04
	452211	Al985513	Hs.233420	ESTs	4.02
25	443292	AK000213	Hs.9196	hypothetical protein	4.01
	420911	U77413	Hs.100293	O-linked N-acetylglucosamine (GlcNAc) tr	4.00 3.95
	428738	NM_000380	Hs.192803	xeroderma pigmentosum, complementation g	
	430456	AA314998	Hs.241503	hypothetical protein	3.95
20	437531	Al400752	Hs.112259	T cell receptor gamma locus	3.93
30	428695	Al355647	Hs.169999	purinergic receptor (family A group 5)	3.91
	410011	AB020641	Hs.57856	PFTAIRE protein kinase 1	3.91
	446494	AA463276	Hs.268906	WW Domain-Containing Gene	3.91
	409928	AL137163	Hs.57549	hypothetical protein dJ473B4	3.90 3.90
25	411598	BE336654	Hs.70937	H3 histone family, member A	3.90
35	425707	AF115402	Hs.11713	E74-like factor 5 (ets domain transcript	3.89
	451806	NM_003729	Hs.27076	RNA 3'-terminal phosphate cyclase	3.89
	401045			1 # F-1 1 DIFFE 5170400	3.89
	437372	AA323968	Hs.283631	hypothetical protein DKFZp547G183	3.88
40	417067	AJ001417 AF102546	Hs.81086 Hs.83931	solute carrier family 22 (extraneuronal dachshund (Drosophila) homolog	3.88
40	410467 431930	AP102546 AB035301	Hs.272211	cadherin 7, type 2	3.88
	453047	AW023798	Hs.286025	ESTs	3.88
	401785	AWU23790	ms.200025	Edia	3.88
	458229	Al929602	Hs.177	phosphatidylinositol glycan, class H	3.86
45	406414	MISESOUE	113.177	pricepriatelymicanol grycent, class 11	3.86
73	412494	AL133900	Hs.792	ADP-ribosviation factor domain protein 1	3.84
	418329	AW247430	Hs.84152	cystathionine-beta-synthase	3.83
	424850	AA151057	Hs.153498	chromosome 18 open reading frame 1	3.82
	427585	D31152	Hs.179729	collagen, type X, alpha 1 (Schmid metaph	3.82
50	423052	M28214	Hs.123072	RAB3B, member RAS oncogene family	3.62
50	416111	AA033813	Hs.79018	chromatin assembly factor 1, subunit A (	3.82
	419423	D26488	Hs.90315	KIAA0007 protein	3.80
	429643	AA455889	Hs.167279	FYVE-finger-containing Rab5 effector pro	3.80
	431499	NM 001514	Hs.258561	general transcription factor IIB	3.80
55	444078	BE246919	Hs.10290	U5 snRNP-specific 40 kDa protein (hPrp8-	3.78
	430291	AV660345	Hs.238128	CGI-49 protein	3.76
	431637	Al879330	Hs.265960	hypothetical protein FLJ10583	3.74
	440411	N30256	Hs.151093	hypothetical protein DKFZp434G1415	3.74
	405917				3.74
60	451230	BE546208	Hs.26090	hypothetical protein FLJ20272	3.73
	429597	NM_003816	Hs.2442	a disintegrin and metalloproteinase doma	3.73
	415075	L27479	Hs.77889	Friedreich ataxia region gene X123	3.72
	440351	AF030933	Hs.7179	RAD1 (S. pombe) homolog	3.70
	443603	BE502601	Hs.134289	ESTs, Weakly similar to KIAA1063 protein	3.70
65	446965	BE242873	Hs.16677	WD repeat domain 15	3.70
	412350	AI659306	Hs.73828	prolein tyrosine phosphatase, non-recept	3.70
	433852	Al378329	Hs.126629	ESTs	3.70
	447397	BE247878	Hs.18442	E-1 enzyme	3.68
	405718				3.68

	425217	AU076696	Hs.155174	CDC5 (cell division cycle 5, S. pombe, h	3.68
	421734	Al318624	Hs.107444	Homo sapiens cDNA FLJ20562 fis, clone KA	3.67
	427221	L15409	Hs.174007	von Hippel Lindau syndrome	3,67
	402408				3,86
5	452946	X95425	Hs.31092	EphA5	3.66
	419078	M93119	Hs.89584	Insulinoma-associated 1	3.66
	427144	X95097	Hs.2126	vascactive intestinal peptide receptor 2	3.65
	423396	Al382555	Hs.127950	bromodomain-containing 1	3.65
	446320	AF126245	Hs.14791	acyl-Coenzyme A dehydrogenase family, me	3.63
10	404939				3.62
	403137				3.60
	437162	AW005505	Hs.5464	thyroid hormone receptor coactivating pr	3.60
	404210				3.59
	443775	AF291664	Hs.204732	matrix metalioproteinase 26	3.56
15	452501	AB037791	Hs.29716	hypothetical protein FLJ10980	3.56
	422443	NM_014707	Hs.116753	histone deacetylase 7B	3.55
	420230	AL034344	Hs.284186	forkhead box C1	3.55
	418428	Y12490	Hs.85092	thyroid hormone receptor interactor 11	3.54
	433002	AF048730	Hs.279906	cyclin T1	3.53
20	405793				3.52
	457940	AL360159	Hs.306517	Homo sapiens TRIpartite motif protein ps	3.52
	402444				3.52
	418250	U29926	Hs.83918	adenosine monophosphate deaminase (isofo	3.51
	414222	AL135173	Hs.878	sorbitol dehydrogenase	3.51
25	422384	AA224077	Hs.42438	Sm protein F	3.50
	447805	AW627932	Hs.19614	gemin4	3.50
	454265	H03556	Hs.300949	ESTs, Weakly similar to thyroid hormone	3.50
	423445	NM_014324	Hs.128749	alpha-methylacyl-CoA racemase	3.48
	413435	X51405	Hs.75360	carboxypeptidase E	3.46
30	447210	AF035269	Hs.17752	phosphatidylserine-specific phospholipas	3.46
	426931	NM_003416	Hs.2076	zinc finger protein 7 (KOX 4, clone HF.1	3.45
	408418	AW963897	Hs.44743	KIAA1435 protein	3.45
	421887	AW161450	Hs.109201	CGI-86 protein	3.44

## Table 7: 42 GENES ENCODING SMALL MOLECULE TARGETS UP-REGULATED IN PROSTATE CANCER COMPARED TO NORMAL ADULT TISSUES

Table 7 shows 42 genes up-regulated in prostate cancer compared to normal adult tissues that are likely to be small molecule targets. These were selected as for Table 5 and the predicted protein contained a structural domain that is indicative of a drugable structure (e.g. protease, kinase, phosphatase, receptor). The functional domain is indicated for each gene.

10 Pkey:

Unique Eos probeset identifier number Exemplar Accession number, Genbank accession number ExAcon:

UnigenelD: Unigene number
Unigene Title: Unigene gene title
PSDomain: Protein Structural Domain 15 R1: Ratio of tumor vs. normal tissue

	Pkey	ExAcon	UnigeneiD	Unigene Title	PSDomain	R1	
20	426747	AA535210	Hs.171995	kallikrein 3, (prostate specific antigen	trypsin	31.80	
		X07730	Hs.171995	kallikrein 3, (prostate specific antigen	trypsin	24.91	
	420757	X78592	Hs.99915	androgen receptor (dihydrotestosterone r	Androgen_recep,hormone_rec,zf-C4	19.72	
	408430	S79876	Hs.44926	dipeptidylpeptidase IV (CD26, adenosine	DPPIV_N_term,Peptidase_S9	16.28	
	430226	BE245562	Hs.2551	adrenergic, beta-2-, receptor, surface	7tm_1	15.40	
25	411096	U80034	Hs.68583	mitochondrial intermediate peptidase	Peptidase_M3	14.81	
	440286	U29589	Hs.7138	cholinergic receptor, muscarinic 3	7tm_1	12.04	
	420381	D50640	Hs.337616	phosphodiesterase 3B, cGMP-inhibited	PDEase	11.10	
	407021	U52077		gb:Human mariner1 transposase gene, comp	SET,Transposase_1	11.02	
	401424				arginase	9.58	
30	410001	AB041036	Hs.57771	kalikrein 11	trypsin	9.03	
	428330		Hs.2256	matrix metalloproteinase 7 (matrilysin,	Peptidase_M10	8.76	
		AF071202		ATP-binding cassette, sub-family C (CFTR	ABC_tran,ABC_membrane	7.64	
		860000fW	Hs.94210	eyes absent (Droscphila) homolog 1	Hydrolase	7.20	
~~		NM_002742	Hs.2891	protein kinase C, mu	pkinase,DAG_PE-bind,PH	6.49	
35		NM_012093		adenylate kinase 5	adenylalekinase	6.00	
		X03635	Hs.1657	estrogen receptor 1	Oest_recep.zf-C4,hormone_rec	5.78	
		AF189723	Hs.106778	ATPase, Ca++ transporting, type 2C, memb	E1-E2_ATPase,Hydrolase	5.37	
		NM_004915		ATP-binding cassette, sub-family G (WHIT	ABC_tran	5.31	
40		M73700		lactotransferrin	transferrin,7tm_1	5.29	
40		X69208	Hs.606	ATPase, Cu++ transporting, alpha polypep	E1-E2_ATPase,Hydrolase,HMA	5.08	
	403047				trypsin	4.91	
		D42063		RAN binding protein 2	Ran_BP1,zf-RanBP,TPR,pro_isomeras		4.88
		NM_001141		arachidonate 15-lipoxygenase, second typ	ipoxygenase,PLAT	4.82 4.82	
45		W15267		low density lipoprotein receptor-related	Idl_recept_b,Idl_recept_a,EGF T4_delodinase	4.82	
43	423740	NM_013989		delodinase, lodothyronine, type II aminopeptidase puromycin sensitive	Peptidase M1	4.24	
		NM 005923		mitogen-activated protein kinase kinase	pkinase	4.21	
		NM_000923		replication factor C (activator 1) 2 (40	AAA,Viral_helicase1	4.20	
		NM_002914 NM_003157	Hs.1087	serine/threonine kinase 2	pkinase _	4.12	
50		Al355647		purinergic receptor (family A group 5)	7tm 1	3.91	
50		AB020641		PFTAIRE protein kinase 1	okinase	3.91	
		AA151057		chromosome 18 open reading frame 1	Idl recept a	3.82	
		AI659306	Hs.73826	protein tyrosine phosphatase, non-recept	Y_phosphatase,Band_41,PDZ	3.70	
		BE247676		E-1 enzyme	Hydrolase	3.68	
55	452946			EphA5	EPH_lbd,fn3,pkinase,SAM	3.66	
-	427144		Hs.2126	vasoactive intestinal peptide receptor 2	7tm 2	3.65	
		AF291664		matrix metalloproteinase 26	Peptidase_M10	3.56	
		AL360159		Homo sapiens TRipartite motif protein ps	SPRY,7tm_1	3.52	
		U29926			A_deaminase	3.51	
60	413435	X51405		carboxypeptidase E	Zn_carbOpept	3.46	
	447210	AF035269	Hs.17752	phosphatidylserine-specific phospholipas	lipase	3.46	

## TABLE 8: 136 GENES SIGNIFICANTLY DOWN-REGULATED IN PROSTATE CANCER COMPARED TO NORMAL PROSTATE

Table 8 shows 136 genes significantly down-regulated in prostate cancer compared to normal prostate. These were selected from 59680 probesets on the Affymetrix/Eos Hu03 GeneChip array such that the ratio of "average" normal prostate to "average" prostate cancer rissues was greater than or equal to 2. The "average" normal prostate level was set to the mean amongst 4 normal prostate tissues. The "average" prostate cancer level was set to the 85th percentile amongst 73 tumor samples. In order to remove gene-specific background levels of non-specific hybridization, the 10th percentile value amonest all the tissues was subtracted from

amongst 7.3 tumor samples. In order to remove gene-specific background reversion from o specific hybridization, the 10th percentile value amongst all the tissues was subtracted from both the numerator and the denominator before the ratio was evaluated.

Unique Eos probeset identifier number

15	ExAcon: Unigene Unigene	ID:	Unigene Unigene	gene title	
	R1:		Ratio of	normal prostate to prostate cancer	
20	Pkey	ExAccn	UnigeneID	Unigene Title	R1
20	425932	M81650	Hs.1968	semenogelin I	57.89
	425545	N98529	Hs.158295	Human mRNA for myosin light chain 3 (MLC	19.70
	426752	X69490	Hs.172004		15.25
	442082	R41823	Hs.7413	ESTs; calsyntenin-2	10.05
25	407245	X90568	Hs.172004		9.38
	422711	D60641		Homo sapiens mRNA; cDNA DKFZp586I1518 (1	9.05
	420813	X51501	Hs.89949	prolactin-induced protein	8.18
	411987	AA375975	Hs.183380	"ESTs, Moderately similar to ALU7_HUMAN	7.45
	404567				5.62
30	416030	H15261	Hs.21948	ESTs	5.51
	444892	Al620617	Hs.148565	ESTs	5.27
	444573	AW043590	Hs.225023	ESTs	5.20
	428068	AW016437	Hs.233462	ESTs	5.08
	437440	AA846804	Hs.123694	ESTs	4.95
35	404113				4.75
				hypothetical protein FLJ13164	4.75
			Hs.188181		4.63
		AV654382	Hs.17947	"ESTs, Weakly similar to K02F3.10 [C.ele	4.53
	405163				4.49
40	405227				4.45
			4Hs.37048	statherin	4.45
			Hs.22968	ESTs	4.40
		U35637		"gb:Human nebulin mRNA, partial cds"	4.03
	403612				4.02
45			Hs.135646		4.00
				Homo sapiens clone TUA8 Crl-du-chat regi	3.98
			Hs.128993	"ESTs, Weakly similar to KIAA0465 protei	3.95
	459367	BE148877		*gb:CM4-HT0244-111199-040-h12 HT0244 Hom	3.95
50				zinc transporter	3.92
50		AW860972		"gb:QV0-CT0387-180300-167-h07 CT0387 Hom	3.85 3.75
			Hs.8/150	Human clone A9A2BR11 (CAC)rl/(GTG)n repea	3.75
		AF089478		*gb:AF089478 Homo saplens astrocytoma li	3.60
	403649			and the first transfer to	3.58
55		H13139		paired-like homeodomain transcription fa	3.50 3.51
33		AA196241		"troponin T1, skeletal, slow" ESTs	3.45
					3.45
	427419	NM_00020	0Hs.177888	mstant 3	3.35
	420777	AA280223	Hs.130865	EOIS ECT-	3.31
60			Hs.161008		3.30
UU		R02018		"Ank, mouse, homolog of" "EST, Highly similar to ubiquitin-protei	3.30
			Hs.14/1/4 Hs.292776		3.26
			Hs.83870		3.16
	400440	X83957	115.000/0	Howaiii	3.10

3.06

413778 AA090235 Hs.75535 \*myosin, light polypeptide 2, regulatory

		AA090235	HS.75535	myosin, ignt polypapida 2, regulatory	3.00
		AW838068		*gb:QV3-LT0048-010300-109-f02 LT0048 Hom	3.05
		AA830811			2.98
-		Al476318	Hs.192480		2.95
5		H00093		*gb:ph8f12u_19/1TV Outward Alu-primed hn	2.92
	405678				2.85
		S73840	Hs.931		2.81
		AW189097			2,78
	433968	AL157518	Hs.90421		2.73
10	438522	AA809431	Hs.258886		2.73
	436562	H71937	Hs.169756		2.68
	412417	AA102268	Hs.42175	ESTs	2.67
	455590	BE072259		"gb:QV4-BT0536-271299-059-g04 BT0536 Horn	2.65
		F07953	Hs.16085	putative G-protein coupled receptor	2.65
15	428729	AL162331		hypothetical protein FLJ10619	2.64
	408537	AW207734		gb:UI-H-BI2-age-h-01-0-UI.st NCI_CGAP_S	2.63
	424706	AA741336	Hs.152108	transcriptional unit N143	2.63
		BE072092		gb:PM4-BT0532-160200-003-b11 BT0532 Hom	2.63
			Hs.929		2.62
20		AA758538			2.60
		Al933794			2.58
		R20723	Hs.124764		2.58
		AA829828			2.52
				"ESTs, Highly similar to FXD3_HUMAN FORK	2.51
25		Al689154			2.50
		AA737400			2.50
	410028	AW576454	Hs 258553	FSTs	2.46
					2.45
		AI638562		"gb:ts50a10.x1 NCL_CGAP_Ut1 Homo saplens	2.44
30		AA015767	He 193587		2.40
50		H87863	Hs.151380		2.36
		AW600293		*gb:EST00049 pGEM-T library Homo sapiens	2.36
	400001	7111000200		AFFX control: BioB-3	2.36
		Z45365		*gb:HSC2NF061 normalized infant brain cD	2.36
35		AW872527	Hs 59761	ESTs	2.36
55	423341	AW242394	Hs.252495	ESTs	2.36
		AA742221			2.35
		AJ002784		gb:Homo sapiens mRNA; fetal brain cDNA 5	2.33
		AA744550	Hs.136345		2.32
40	401974				2.31
		AL044498	Hs.133262	*ESTs, Weakly similar to PH0217 roverse	2.31
				transient receptor potential channel 5	2.25
		AI949371			2.25
		R15337		"Homo sapiens cDNA FLJ10532 fis, clone N	2.25
45		Al762250		ESTs	2.24
	405420				2.23
		AW851258		"qb: L3-CT0220-160200-066-H06 CT0220 Hom	2.23
	438224	AA933999		"gb:on91f04.s1 Soares_NFL_T_GBC_S1 Homo	2.23
		BE008347		"gb:CMO-BN0154-080400-325-h04 BN0154 Hom	2.23
50		BE252470		"qb:601108292F1 NIH_MGC_16 Homo sapiens	2.23
	437010	AA741368	Hs.291434	ESTs	2.23
		Al914279			2.22
	403375				2.21
	455060	AW853441		"ab:RC1-CT0252-030100-023-g09 CT0252 Hom	2.21
55	409792	AW854153		*gb:RC3-CT0254-060400-029-d03 CT0254 Hom	2.20
	421154	AA284333	Hs.287631	"Homo sapiens cDNA FLJ14269 fis, clone P	2.19
	401963				2.18
	435034	AF168711	Hs.159397	x 010 protein	2.18
				K/AA0553 protein	2.18
60		AW297599			2.17
		Al733395			2.17
		AA236233			2.18
		H91800	Hs.124156		2.16
		R54109	Hs.26096		2.18
65		AA989835			2.15
		Al133482			2.15
		AA425562		*gb:zw46e05.r1 Soares_total_fetus_Nb2HF8	2.15
	437101	AA744518	Hs.120610	ESTs	2.15
	428793	AC004957	Hs.298975	*ESTs, Highly similar to collapsin-2-lik	2.15
				170	

	415708 459619	H56475		*gb:yt87d11.r1 Soares_pineal_gland_N3HPG	2.13
		AK000134	Hs 179100	hypothetical protein FLJ20127	2.12
			Hs.184354		2.10
5		AW809157	110.101001	"ab:RC0-ST0118-041099-031-c07_1 ST0118 Homo sapiens cDNA, mRNA sequence"	2.10
5	403087	/11000107		Parion of the parions on and a series and a	2.10
	403869				2.10
		D81194	Hs.282499	EQTe	2.10
		H29505	110.202400	"ab:vm60d10.r1 Scares infant brain 1NIB Homo saplens cDNA clone 5", mRNA sequence"	2.10
10		H11257	Hs.295233		2.09
10		BE218221	Hs.190044		2.08
		BE274360	NS. 180044	"gb:601121038F1 NIH_MGC_20 Homo saplens cDNA clone 5', mRNA sequence"	2.08
	405455	DE274000		go do i iz 1000r i ivii _mao_zo i iviio sapistis obrita ciotto o i ilitara sequence	2.08
		AA332652		"ab:EST36627 Embryo, 8 week I Homo saplens cDNA 5" end similar to similar to	2.00
15	423043	AVA332002		monoamine oxidase B. mRNA sequence"	2.08
13	406135			monoamme oxidase o, minivi sequence	2.07
		DE0 40400	Hs.121385	ECTA	2.07
		BE246180	HS, 12 1385	ESIS	2.05
	403493	*1000007	11. 070404	STOTE WAS ALLEGADING THE STREET ALLEGADES AND ALLEGADES AN	2.00
20	444514	Al682905	H8.270431	"ESTs, Weakly similar to ALU1_HUMAN ALU SUBFAMILY J SEQUENCE	2.05
20				CONTAMINATION WARNING ENTRY [H.saplens]*	
			Hs.192868		2.05
		AB020695	Hs.91662	KIAA0888 protein	2.03
	405900				2.03
		AW974438	Hs.194136	"ESTs, Moderately similar to AF091457 1 zinc finger protein RIN ZF [R.norvegicus]"	2.02
25	400007			AFFX control: BloDn-5	2.01
	406978	M64358		"gb:Human rhom-3 gene, exon."	2.00

TABLE 8A shows the accession numbers for those primekeys lacking a unigeneID in Table 8. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (DoubleTwist, Oakland California). The Genbank accession numbers for sequences comprising each cluster are listed in the "Accession" column.

Unique Eos probeset identifier number Gene cluster number

5

10

Pkey: CAT number:

	Accession:		Genbank accession numbers
15	Pkey	CAT number	Accessions
	407764	1014849 1	BE008347 BE008320 BE083307 BE083311 AW075988
	403537	1064753_1	AW207734 D60164 D81150 D81078 D61356 AW996804
	409792	1154677_1	AW854153 AW500210 BE145772 AW501310
20	410881		AW809157 AW812181 AW812175 AW812172 AW812161 AW812165
	411762	1256906_1	AW860972 AW862598 AW862599 AW860988 AW860983 AW860989 AW860925 AW860922 AW860986 AW860984 AW860989
		1353792 1	BE072092 BE072106 BE072096 BE072098 BE072103
	413549	1375933 2	BE252470 BE147573
	415708	1548209_1	H56476 P29401 F34552
25			Z45365 R25905 H05203 T77496
			Al638562 T16929 H13401 F07773 R55836
			AW838068 AW837986 AW838067 AA322487 AW837936
	423843	232510 1	AA332652 AA331633 AW999369 AW902993 BE170475 AA378845 AW964175 AI475221
			AA425582 AI860208 AA346646 N22655 AW811775 AW811786
30		274259 -1	BE274380
		347718 2	H00093 H00079 H00070 H00054 H00049 H00063 AW905906 AW905241 AW905410 AW905307 AW905411 AW905240
	AW9052		
			AW905352 AW905304 AW905239 AW905242 AW905243 H00087
	438224	452656_1	AA933999 AA781181
35		740749_1	H29505 R18575 Z43580 T48738 Al435454 BE004683
		863269_1	AW800293 AI767468
			AW851258 AW851435 AW851106 AW351421
		1251259 1	AW853441 BE145228 BE145218 BE145162 BE145283
	455590		BE072259 BE072230 BE007911
40	459311	543550 1	AF069478 AF069479 AF069480

TABLE 8B shows the genomic positioning for those primekeys lacking unigene ID's and accession numbers in table 8. For each predicted exon, we have listed the genomic sequence source used for prediction. Nucleotide locations of each predicted exon are also listed.

5							
10	Pkey: Ref;		Sequence sour publication entil	corresponding to an Eos proboset co. The 7 digit numbers in this column are Genbank Identilliar (GI) numbers. "Dunham I. et al." refors to the lled "The DNA man chromosome 22." Dunham I. et al., Nature (1999) 402-499-495.			
10	Strand: Nt_position	n:	Indicates ON A stand from which exons were predicted.  Indicates ONA stand from which exons were predicted.  Indicates nucleotide positions of predicted exons.				
15	Pkey	Ref	Strand	Nt_position			
	401963	3126783	Plus	51382-51521			
	401974	3126777	Plus	85330-85683			
	403037	8954241	Plus	169511-169795			
20	403375	9255944	Minus	92554-92795			
	403493	7341425	Plus	157568-159084			
	403612	8469060	Minus	94723-94859			
	403649	8705159	Minus	27141-27247			
	403869	7280046	Minus	34379-34583			
25	404113	9588571	Minus	13446-13646			
	404567	7249169	Minus	101320-101501			
	405163	9968267	Minus	161171-161299			
	405227	8731245	Minus	22550-22802			
	405420	7211837	Minus	13428-13582			
30	405455	7656675	Plus	134112-134671			
	405678	4079670	Plus	151821-152027			
	405900	6758795	Minus	71181-71535			
	406135	9164918	Minus	65489-65715			

## TABLE 9: 1001 GENES SIGNIFICANTLY UP-REGULATED IN NORMAL PROSTATE COMPATED TO PROSTATE CANCER

Table 9 shows 1001 genes significantly up-regulated in prostate cancer compared to normal 5 prostate. These were selected from 59680 probesets on the Affymetrix/Eos Hu03 GeneChip array such that the ratio of "average" normal prostate to "average" prostate cancer tissues was greater than or equal to 8.14. The "average" normal prostate level was set to the mean amongst 4 normal prostate issues. The "average" prostate cancer level was set to the Stsh percentile amongst 73 tumor samples. In order to remove gene-specific background levels of 10 non-specific hybridization, the 10<sup>th</sup> percentile value amongst all the tissues was subtracted from both the numerator and the denominator before the ratio was evaluated.

15	Pkey: ExAcon: Unigene Unigene R1:	iD:	Exemplar A Unigene nu Unigene ge			
20	Pkey	ExAccn	UnigenelD	Unigene Title	R1	
20	451002	AA013299	Hs.8018	ESTs, Weakly similar to ALU3_HUMAN ALU S	1684.00	
		AA689465	Hs 188999		738.00	
		Al078027	Hs.169338	ESTs	246.86	
		AA928118	Hs.272065		245.20	
25		AK000185		gb:Homo saplens cDNA FLJ20178 fis, clone	222.00	
	405932	181000100		,	221.33	
		AA864330	Hs.166520	ESTs	212.00	
		AI686550	Hs.174481	ESTs	163.20	
		AJ474866	Hs.193237	ESTs	149.45	
30		NM_002118	Hs.1162	major histocompatibility complex, class	126.11	
		M36860	Hs.9295	elastin (supravalvular aortic stenosis,	123.27	
		AW138330	Hs.233778		120.00	
	418917	X02994	Hs.1217	adenosine deaminase	106.75	
	404407				105.71	
35		AI652926	Hs.128395	ESTs	100.53	
		AA608684	Hs.121705	ESTs, Moderately similar to ALUC_HUMAN !	94.00	
		U83527		qb:HSU83527 Human fetal brain (M.Lovett)	89.18	
		F06495		gb:HSC1AB051 normalized infant brain cDN	87.73	
	424239	M67439	Hs.143526	dopamine receptor D5	88.62	
40	444143	AW747996	Hs.160999	ESTs	86.43	
	401672				77.26	
	430590	AW383947	Hs.246381	CD68 antigen	68.47	
	411972	BE074959		gb:PM0-BT0582-310100-001-f08 BT0582 Homo	68.00	
	448992	AI766053	Hs.188346	ESTs	61.26	
45	408828	BE540279		gb:601059857F1 NIH_MGC_10 Homo sapiens C	57.71	
		AW451693	Hs.220826	ESTs	56.40	
	402964				54.67	
	422673	N59027		gb:yv59d11.r1 Soares fetal liver spleen	54.00	
		AA372275		Homo sapiens cDNA FLJ11393 fis, clone HE	54.00	
50	438907	R32704	Hs.301296	ESTs	52.96	
	405172				52.96	
		AW137088	Hs.144857		52.32	
		AW592931	Hs.256298		51.63	
		AB028989		mitogen-activated protein kinase 8 inter	50.98	
55		AA703679	Hs.106999	ESTs, Weakly similar to SYT5_HUMAN SYNAP	49.60	
		AA339666		gb:EST44776 Fetal brain I Homo saplens c	48.90	
		T54095		gb:ya92c05.s1 Stratagene placenta (93722	47.98	
		AA424163	Hs.156895		46.83	
		Al700148	Hs.283626		43.57	
60		AA485224		G protein-coupled receptor kinase-Intera	43.00	
		AA837098	Hs.269933		42.70	
	438854	AF074994	Hs.24240	ESIS	42.67	

	406134				42.43
	457319	AA480895	Hs 201552	ESTs, Weakly similar to T17288 hypotheti	42.31
		AA070266		gb;zm89d04.r1 Stratagene neuroepithelium	42.25
	401124	7010/0200		go,zarazdo4.11 Gadasgene necroeprimenani	41.61
~					
5		Al371157	Hs.178538		40.00
	420317	AB006628	Hs.96485	KIAA0290 protein	39.64
	457586	AW062439		gb:MR0-CT0060-120899-001-f08 CT0060 Homo	39.60
		AA923278	He gangns	ESTs, Weakly similar to protease [H.sapi	38.73
			Hs.178364		38.06
10		BE221682			
10		W79114	Hs.58558	ESIS	36.69
	433686	AA604799	Hs.136528	ESTs, Moderately similar to ALU1_HUMAN A	35.29
	417993	AW963705	Hs 295806	ESTs, Weakly similar to ALU7_HUMAN ALU S	36.18
		AA936282	Hs.120397		36,10
		AA333990		coagulation factor XIII, A1 polypeptide	36.08
15					
15		BE314852		hypothetical protein FLJ10257	36.00
	415911		Hs.124952	ESTs	36.00
	457502	AA076049	Hs.274415	Homo sapiens cDNA FLJ10229 fis, clone HE	35.23
	421566	NM_000399	Hs.1395	early growth response 2 (Krox-20 (Drosop	35.20
	401468	rem_ooccoo	11011000	can't Brown to the same of the same	34.89
20		41000450	U- 01440E	FOT-	34.60
20		Al220150	Hs.211195		
	433801	BE350738	Hs.123993	ESTs, Weakly similar to T00366 hypotheti	33.24
	454977	AW848032		ab:lL3-CT0214-231299-053-D11 CT0214 Homo	32.96
	402828			•	32.93
		AW518944	Un 70000	Homo sapiens cDNA: FLJ23125 fis, clone L	31.76
25		WALD 10044	115.7 0020	Homo sapiena contra i cazarza na, cione c	31.68
25	402842				
		AA285363		gb:HTH280 HTCDL1 Homo sapiens cDNA 51/31	31.59
	401631	F05183	Hs.1799	CD1D antigen, d polypeptide	31.26
	408057	AW139565		gb:UI-H-BI1-aea-d-04-0-UI.s1 NCI_CGAP_Su	31.24
		H81795		gb:ys68a10.r1 Soares retina N2b4HR Homo	31.20
20			Hs.291797		31.09
30	438694				31.05
		AF103907	Hs.171353	prostate cancer antigen 3	29.78
	428796	AU076734	Hs.193665	solute carrier family 28 (sodium-ccupled	29.76
	452549	Al907039		gb:PM-BT134-020499-566 BT134 Homo sapien	29.59
		BE244074	Hs 285531	regulator of Fas-Induced apoptosis	29.53
35		AI870175	Hs.13957	ECTo	29.47
33					29.22
		R07566	Hs.73817		
	459081	W07808		gb:zb03a12.r1 Soares_fetal_lung_NbHL19W	29.20
	448702	AW102670	Hs.122464	ESTs	29.13
		U80456	Hs.27311	single-minded (Drosophila) homolog 2	28.74
40		W84893	Hs.9305		28.61
-10	457004	AD00000	H=042004	VIA A1007 protoin	28.24
	40/324	ADU2099U	115.243901	NAMIOS/ protein	
	424247	X14008	HS.234734	KIAA1087 protein lysozyme (renal amyloidosis) ESTs alpha-methylacyt-CoA racemase	28.18
	457140	Al279960	Hs.178140	ESTs	28.12
	444151	AW972917	Hs.128749	alpha-methylacyl-CoA racemase	28.06
45	457669	AW104257	He 123426	ESTs, Wealdy similar to putative serine/	27.61
.,,		AV650262		GRO2 oncogene	27.36
		AVOUCEUZ	115.73700	dhoz dhogene	27.33
	405495				
	406516				27.25
	407997	AW135429	Hs.243577	ESTs	26.96
50	442115	AW135429 AW452332	Hs.257554	ESTs	26,36
	100038	T97490		small inducible cytokine subfamily A (Cy	26.34
		107400	110,000	ornal madding of terms cantainly it (o)	26.32
	402838				- 26.21
		AI979284	Hs.200552		
	417153	X57010	Hs.81343	collagen, type II, alpha 1 (primary oste	26.20
55	439792	NM_014856	Hs.6684	KIAA0476 gene product	25.91
		Al682088			25.60
		AL133660	Un 142022	Homo sapiens mRNA; cDNA DKFZp434M0927 (f	25.57
					25.57
		BE391090	Hs.280278	ES1	
		NM_005188		Cas-Br-M (murine) ecotropic retroviral t	25,48
60	424778	AA251048	Hs.153042	lymphocyte antigen 9	25.42
		AA063426		gb:zf70c08.s1 Soares_pineal_gland_N3HPG	25.25
		AW083491	Hs.31196		25.22
				gb:51f10 Human retina cDNA randomly prim	25.01
	418392	W28573	11-0505	COT- 11 -14 -imites to CO   I   I   I   I   I	
		T74588	Hs.8509	ESTs, Weakly similar to CO3_HUMAN COMPLE	24.85
65	422940	BE077458		gb:RC1-BT0606-090500-015-b04 BT0606 Homo	24.76
	437571	AA760894	Hs.153023	ESTs	24.74
	433973	Al014723	Hs.131770		24.57
		BE019557	He 11900	Human DNA sequence from clone RP4-583P15	24.53
			U- 105700	secreted frizzled-related protein 4	24.49
	421002	AF026692	16,100/00	services intraceriorates bioresis 4	2440

		U25758	Hs.134584		24.49
		AL035588		MyoD family inhibitor	24.10 24.04
		AA357001 AL122081	Hs.34045	hypothetical protein FLJ20764 cadherin related 23	24.00
5		AI208611	Hs.12066		23.89
-		AA215872		gb;zr96e09.s1 NCL_CGAP_GCB1 Homo sapiens	23.83
	406583	AW449674	Hs.47359	ESTs	23.73
		AP204231	Hs.182982	golgin-67	23.62
10		AA136301		gb:zk93g04.s1 Soares_pregnant_uterus_NbH	23.39 23.20
10		NM_001327 AF123050	Hs.167379 Hs.44532	cancer/testis antigen diubiquitin	22.68
		BE243877	Hs.76941	ATPase, Na+/K+ transporting, beta 3 poly	22.65
	418299	AA279530	Hs.83968	integrin, beta 2 (antigen CD18 (p95), ly	22.38
		R68651	Hs.144997		22.26
15		BE367335	Hs.283713		22.08
	415788	AW628686	Hs.78851	KIAA0217 protein	22.04
		AW809637	11.000	gb:MR4-ST0124-261099-015-b07 ST0124 Homo	22.00 21.95
		AI431708 AV653846	Hs.820	homeo box C6 Homo sapiens Chromosome 16 BAC clone CIT	21.94
20		BE071874	113.120201	gb:RC2-BT0522-120200-014-a06 BT0522 Homo	21.84
20	406748		Hs.47431	spectrin, beta, erythrocytic (includes s	21.26
		H14487		gb:ym18c10.r1 Soares infant brain 1NIB H	21.24
	440474	AI207936	Hs.7195	gamma-aminobutyric acid (GABA) A recepto	21.14
25		AI623698		Homo sapiens cDNA: FLJ23529 fis, clone L	21.11 21.10
25	426793	X89887	Hs.172350	HIR (histone cell cycle regulation defec gb:UI-HF-BR0p-ajr-e-05-0-UI.r1 NIH_MGC_5	21.07
	405685	AW502139		gb.ornr-bnop-sp-e-co-d-st.11 Nin_wdc_s	20.90
		AI983207	Hs 192481	ESTs, Weakly similar to SYPH_HUMAN SYNAP	20.84
		AA321355	Hs.285401	ESTs	20.74
30		AW403724	Hs.140	immunoglobulin heavy constant gamma 3 (G	20.73
	401201				20.73
		W28912	Hs.129019	ESTs gb:yr86d10.r1 Sceres fetal liver spleen	20.68 20.67
		H66948 H42679	Hs.77522		20.66
35	400926	1142070	110.71022	major muscompaning complex; ease	20.88
55		NM 004197	Hs.444	serine/threonine kinase 19	20.64
		AW500221	Hs.43616	Homo saplens mRNA for FLJ00029 protein,	20.81
		X60992	Hs.81226	CD6 antigen	20.61
40	405777	*********	H- cocoo	Homo sapiens cDNA FLJ12702 fis, clone NT	20.51
40	424123	AW966158 X58288		protein tyrosine phosphatase, receptor t	20.10
		BE568568	Hs.195704		19.98
	421064	AI245432		tumor necrosis factor, a/pha-induced pro	19.98
	418819	AA228776	Hs.191721	ESTs	19.94
45		AA584854		gb:ne09h11.s1 NCI_CGAP_Phe1 Homo sapiens	19.90 19.84
	404426	U43143	Un 7/0/0	fms-related tyrosine kinase 4	19,79
		NM_012211		integrin, alcha 11	19.62
		NM_006732	Hs.75678		19.57
50	418994	AA296520	Hs.89546		19.56
		AW090198	Hs.4779	KIAA1150 protein	19.52
		AA156781	Hs.83992		19.44
		AL138201 X15675	Hs.82120	Human pTR7 mRNA for repetitive sequence	19.22
55		AW449808		glucosamine (N-acetyl)-6-sulfatase (Sanf	19.21
55	456557	AA284477	Hs.96618		18.77
	440606	AI247422	Hs.129986		18.76
	439845	AL355743		Homo sapiens EST from clone 41214, full	18.85
60		AI807264 AA769062	Hs.205442 Hs.16029	ESTs, Weakly similar to AF117610 1 Inner ESTs, Weakly similar to alternatively sp	18.64 18.62
oo		AA769062 AW043951	Hs.38449	ESTs weakly similar to alternatively sp	18,59
	418329	AW247430	Hs.84152		18.58
	424537	AI673027	Hs.143271	ESTs	18.55
	447742	AF113925		caspase recruitment domain 4	18.52
65	415251		Hs.7124	ESTs	18.47
	440770	AA912815 AI085846	Hs.222078 Hs.25522		18.40 18.32
		U51166		thymine-DNA glycosylase	18.28
		AW501751	Hs.279733		18.15

	417240	N57568	Hs.176028	EST	18.1
		AF229178		leucine rich repeat and death domain con	18.1
	436896	AW977385	Hs.278615	ESTs	18.1
		N90666	He 276770	CDW52 antigen (CAMPATH-1 antigen)	17.9
-					17.8
5		Al971131		ESTs, Weakly similar to alternatively sp	
	429984	AL050102	Hs.227209	DKFZP586F1019 protein	17.8
		AI889114	Hs.195663		17.7
	433867	AK000598	Hs.3618	hippocalcin-like 1	17.7
	421725	AW977724	Hs.75968	thymosin, beta 4, X chromosome	17.7
10		MIGHTER	110.73000	ulymoan, bela 4, A dilloridocino	
10	401515				17.6
	444045	AI097439	Hs.135548	ESTs	17.5
		AL045825	Hs.210197		17.5
	426559	AB001914	Hs.170414	paired basic amino acid cleaving system	17.5
	49941E	T16971	Hs.269014	FSTe	17.5
10					17.5
15		Al188225	Hs.127462		
	432516	R08003	Hs.188013	ESTs	17.4
		AA152106	Hs.4859	cyclin L ania-6a	17.3
			H5.4008		
	414989	T81668		gb:yd29c04.r1 Soares fetal liver spieen	17.3
	444880	AW118683	Hs.154150	FSTs	17.3
20	444000	744110000			17.2
20	417651	R06874	Hs.268628		
	453457	AL037103	Hs.270599	ESTs, Weakly similar to unnamed protein	17.2
		AW452533	Hs.143604	Kaina	17.2
	419078	M93119	Hs.89584		17.1
		BE241624	Hs.82401	CD69 antigen (p60, early T-cell activati	17.1
25					17.1
23		AF003522	HS.250000	delta (Drosophila)-like 1	
	455254	AW877015		gb:QV2-PT0010-250300-096-f12 PT0010 Homo	17.1
		U66468	He 150595	cell growth regulatory with EF-hand doma	17.1
	426678	H08170	Hs.113755	ESIS	17.1
	100103	NM 000361	Hs.2030	thrombomodulin	17.0
30					17.0
30		AB032959		KIAA1133 protein	
	438867	AW451157	Hs.181157	ESTs	16.9
		AA830664	Hs.143974	ESTe	16.9
			118.140974		16.9
	459234	AI940425		gb:CM0-CT0052-150799-024-c04 CT0052 Homo	
	404756				16.9
35		140044	H- 440000	solute carrier family 1 (high affinity a	16.9
22		U18244			
	420568	F09247	Hs.167399	protocadherin albha 5	16.8
		A1076765	Hs.269899	FSTe	16.8
					16.7
	438703	AI803373	Hs.31599	ESTs	
	411424	AW845985		ab:RC2-CT0163-200999-002-H08 CT0163 Homo	16.7
40		7111010000		8	16.8
40	402895				
	422538	NM_006441	Hs.118131	5,10-methenylletrahydrofolate synthetase	16.5
	A47100	AW449802	He 217953	ESTs, Moderately similar to NK-TUMOR REC	16.6
				de this contract Out I bear all the	16.5
		AB002367	Hs.21355	doublecortin and CaM kinase-like 1	
	438587	AW451955	Hs.153065	ESTs	16.5
45	407044	AW190902	He 40008	cysteine knot superfamily 1, BMP entagon	16.5
43					
	410721	R23534	Hs.2730	heterogeneous nuclear ribonucleoprotein	18.5
	437133	AB018319	Hs.5460	KIAA0776 protein	16.4
			110.0400		16.3
		AA047854		gb:zf49g04.r1 Soares retina N2b4HR Homo	
	417315	AI080042	Hs.180450	ribosomal protein S24	16.3
50		AA534908	Hs.2860	POU domain, class 5, transcription facto	16.2
20					16.2
	439882	AA847856	Hs.124565		
	418277	AW135221	Hs.130812	FSTs	16.0
		AW796342		gb:PM2-UM0027-230200-002-h02 UM0027 Homo	16.0
				gu.r me-umoue r-esucus-uuz-nuz umuuz munu	
	420120	AL049610	Hs.95243	transcription elongation factor A (Sif)-	16.0
55		NM 003816	Hs.2442	a disintegrin and metalloproleinase doma	16.0
55					16.0
		AI357412		EST - not in UniGene	
	421684	BE281591	Hs.106768	hypothetical protein FLJ10511	15.9
	400000	AA055800	Hs.222933		15.9
	400099	MODOUM			
	446012	AV656098	HS.172382	hypothetical protein FLJ20001	15.8
60		AA076769		gb:7B02B10 Chromosome 7 Fetal Brain cDNA	15.8
00		3103		0-11-12-11-11-11-11-11-11-11-11-11-11-11-	. 15.8
	405934				
	426108	AA622037	Hs.166468	programmed cell death 5	15.8
	410200	AW291168	Hs.41295	ESTs	15.4
	410208	M11201 100			
	410708	AA534370		Homo sapiens cDNA: FLJ22756 fis, clone K	15.4
65	447342	Al199268	Hs.19322	ESTs; Weakly similar to !!!! ALU SUBFAM!	15.3
00	441042	11100200		gb:CM0-ST0081-130999-054-d02 ST0081 Homo	15.
		AW807530			
	411507	AW850140		gb:lL3-CT0219-261099-023-D11 CT0219 Homo	15.3
		Al916685	Hs.194601		15.2
					15.3
	416292	AA179233	Hs.42390	nasopharyngeal carcinoma susceptibility	15.3

	406638	M13861		gb:Human T-cell receptor active beta-cha	15.26
	446686	AW138043	Hs.156307	ESTs	15.25
		Al623511	Hs.118567		15.24
	444400	AW292830	Hs.255609		15.22
5					
- 3		BE147740	Hs.104558		15.22
		BE244854	Hs.159578	Homo sapiens mRNA for FLJ00020 protein,	15.16
	420748	AA279956	Hs.88672	ESTs	15.14
		AA410506		H.sapiens mRNA for ribosomal protein L18	15.14
		AB023185		calcium/calmodulin-dependent protein kin	15.12
10					
10		Al662096	Hs.50640	ESTs	15.12
	437495	BE177778		gb:RC1-HT0598-310300-012-f07 HT0598 Homo	15.12
	445487	Al239832	Hs.15617	ESTs, Weakly similar to ALU4 HUMAN ALU S	15.06
		AW006783	Hs.6686	ESTs	15.03
	402812	A11000700	1 10.0000	LOIS	15.02
10					
15		AA732480	Hs.293581	ESTs	15.00
	400991				15.00
	415752	BE314524	Hs.78776	Human putative transmembrane protein (nm	14.96
		AA460421	Hs.30875	ESTs	14.90
	403683	791100161	110,00070	2013	14.84
20					
20		NM_004293		guanine deaminase	14.80
	451952	AL120173	Hs.301663		14.72
	424687	.105070	Hs.151738	matrix metalloproteinase 9 (gelatinase B	14.69
		BE617135		gb:601441677F1 NIH_MGC_65 Homo sapiens c	14.67
		AB021225	H- scores	matrix metalloproteinase 17 (membrane-in	14.65
0.5					
25		Al638449	Hs.173031		14.63
	431089	BE041395	Hs.283676	ESTs, Weakly similar to unknown protein	14.60
	459145	AI903354		gb:RC-BT029-100199-117 BT029 Homo sapien	14.55
		AF055575	He 207047	ESTs, Moderately similar to calcium chan	14.54
		MI 000070	110.201041	2018, Moderatory Silisar to Caldull Chair	
00	400952		*		14.48
30		At734009		EST cluster (not in UniGene)	14.44
	407938	AA905097	Hs.85050	phospholamban	14.42
	431676	Al685464	Hs.292638	FSTs	14.40
		AA311443		Homo sapiens mRNA; cDNA DKFZp586E2317 (f	14.36
		AB023199	Hs.27207		14.36
25					
35		AA126419	Hs.301632		14.32
	412368	AW945992	Hs.181125	immunoglobulin lambda locus	14.31
	409055	AW304028	Hs.300578	ESTs	14.23
		W57550		Homo sapiens cDNA FLJ13181 fis, clone NT	14.22
		AL049278			14.22
40				Homo sapiens mRNA; cDNA DKFZp564I153 (fr	
40		BE242639		ubiquitin associated protein	14.22
	421913	AI934365	Hs.109439	osteoglycin (osteoInductive factor, mime	14.22
	452712	AW838616		gb:RC5-LT0054-140200-013-D01 LT0054 Homo	14.22
		AW503398	Hs.210047		14.16
		Y14443	Hs.88219		14.14
45					
45	424909			cell division cycle 25B	14.07
	434078	AW880709	Hs.283683	EST	14.07
	415254	AI815831	Hs.184378	ESTs	14.05
		AI745649	Hs.26549		14.02
	410020		Hs.728	ribonuclease, RNase A family, 2 (liver,	13.98
50				DAG -Od	
50		NM_002890	Hs.758	RAS p21 protein activator (GTPase activa	13.98
	429848	AF145439	Hs.225946	chemokine (C-C motif) receptor 9	13.95
	413729	BE159999		gb:QV1-HT0412-270300-123-d10 HT0412 Homo	13.90
	400125			•	- 13.88
		AW406289	Hs.96593	hypothetical protein	13.85
55					
22		A1479094	Hs.170786		13.80
	422695	AA315158		gb:EST186956 HCC cell line (matastasis t	13.80
	424565	AW102723	Hs.75295	guanylate cyclase 1, soluble, alpha 3	13.78
		H30340		Homo sapiens cDNA: FLJ22050 lls, clone H	13.78
	400004	AI935400	Hs.217286		13.76
60			ns.41/286		
60		AW860158		gb:RC0-CT0379-290100-032-b04 CT0379 Homo	13.75
	410889	X91662	Hs.66744	twist (Drosophila) homolog (acrocephalos	13.74
		A1908236		gb:IL-BT166-180399-010 BT166 Homo sapien	13.72
		AW857913		gb:RC0-CT0323-231199-031-b05 CT0323 Homo	13.69
			11- 400 450		
65		AW015238	Hs.128453		13.67
65		AA365752	Hs.155965	E818	13.62
	402359				13.60
	401044				13.53
		AW502498	He 157150	ESTs. Weakly similar to zinc finger prot	13.53
					13.49
	423090	AA329648	Hs.23804	Lore	10.49

		Al690234		ESTs, Weakly similar to reverse transcri	13.47
		AW578849		ESTs, Weakly similar to unnamed protein	13.46
		AW080339	Hs.211911		13.44
5		AI573283	Hs.38458	gb:yt87c03.r1 Soares_pineal_gland_N3HPG	13.44 13.43
3	402788	H56389		go:yte/cus.ri soares_prieat_gland_names	13.43
		AA886446	Hs.146278	ESTs	13,40
	405411	701000110	Harriotio	2010	13.38
		AW183574	Hs.24218	ESTs	13.34
10		AA132818		ESTs, Weakly similar to coded for by C.	13.33
		AL043004		Human serine/threonine kinase mRNA, pert	13.32
		AI074149	Hs.150905	ESTs, Weakly similar to chondroitin 4-su	13.32
	403838				13.32
15		Z46223		Fc fragment of IgG, low affinity IIIb, r	13.30 13.28
15		AW207552 N41359	Hs.218107	ESTs, Weakly similar to dJ134E15.1 [H.sa	13.28
		AW451101		ESTs, Moderately similar to hexokinase I	13.27
		AF043722	Hs.99491	RAS guanyl releasing protein 2 (calcium	13.26
	420052		Hs.44410	ESTS	13.25
20		NM 002984	Hs.75703		13.25
	403851				13.24
		W07492	Hs.157101	ESTs	13.21
		AI762836		ESTs, Moderately similar to ALU2_HUMAN A	13.21
05		AB033113	Hs.50187		13.20 13.19
25		R21968	Hs.248746	G protein-coupled receptor kinase-Intera	13.17
		BE386844 Al796320		Homo sapiens cDNA FLJ13545 fis, clone PL	13.16
		AA278362		Homo sapiens cDNA FLJ12334 fis, clone MA	13.14
		BE262802	Hs.4909	dickkopf (Xenopus laevis) homolog 3	13.07
30		NM_001621		aryl hydrocarbon receptor	13.06
	414789	AA155859	Hs.79708		13.05
		BE387790	Hs.26369	ESTs	13.04
		T99719		Homo sapiens cDNA: FLJ22389 fis, clone H	13.03
35		AW964806		ESTs, Weakly similar to putative glycine	13.02 13.00
33		Al660552 H20276	Hs.31742	ESTs, Weakly similar to A56154 Abī subst ESTs	13.00
		AL137466	Hs.97277		12.99
		N75276	Hs.135904		12.98
		AA032197	Hs.102558		12.96
40	419953	BE267154	Hs.125752		12.98
		NM_004354	Hs.79069		12.94
		AA015879	Hs.33536	ESTs	12.93
		AW903830		gb:CM4-NN1037-250400-155-h04 NN1037 Homo	12.93 12.92
45		AW161319 D63480	Hs.12915	ESTs KIAA0146 protein	12.92
45		NM 001259	Hs.38481		12.92
		AA534163	Hs.5476	serine protease Inhibitor, Kazal type, 5	12.90
		H41324	Hs.31581		12.88
		D63216		frizzled-related protein	12.88
50		AU076649	Hs.76556	growth arrest and DNA-damage inducible 3	12.88
		AA587775	Hs.66295	Homo sapiens HSPC311 mRNA, partial cds	12.84
		BE077084		gb:RC5-BT0603-220200-013-C07 BT0603 Homo	12.84
		NM_000378	Hs.75596	interleukin 2 receptor, beta	- 12.80 12.80
55		BE167229 BE265839	Hs.29206 Hs.12126	Homo sapiens clone 24659 mRNA sequence hepatocellular carcinoma-associated anti	12.78
33		U97018	Hs.12451	echinoderm microtubule-associated protei	12.78
		W26786	110112-101	gb:15o7 Human retina cDNA randomly prime	12.77
		AU076643	Hs.313	secreted phosphoprotein 1 (osteopontin,	12.76
	447769	AW873704	Hs.48764	ESTs	12.76
60		Al306389	Hs.76240	adenylate kinase 1	12.76
		D83407	Hs.156007	Down syndrome critical region gene 1-lik	12.68
		H85157	Hs.40696	ESTs	12.66
	405856	DEGGTOAR	No 75004	t drulla macifia chanazana a	12.66 12.65
65	412935	BE267045	Hs.75064	tubulin-specific chaperone c	12.62
U.J		AA889120	Hs.110697	Homeo box A10	12.62
		NM_001454		forkhead box J1	12.62
	403137				12.60
	430226	BE245562	Hs.2551	adrenergic, beta-2-, receptor, surface	12.57

		AJ133123	Hs.20196	adenylate cyclase 9	12.56
		F07097	Hs.300628	Homo sapiens mRNA full length insert cDN	12.54
	405238				12.52
5		AA071051		gb:zm58e05.s1 Stratagene fibroblast (937	12.47
,		AA767689	Hs.10242	ESTs ESTs	12.47
		AW978731 Al039201	Hs.301824 Hs.54548		12.44
		BE077546	Hs.31447	ESTs	12.42
		AW984111	FIS.01447	ab:RC0-HN0007-160300-011-109 HN0007 Homo	12.42 12.40
10		AI926047	Hs.162859		12.37
10		M36564	Hs.64016		12.36
		R96698	Hs.35598	ESTs	12.36
		AW189232	Hs.39140		12.36
	432892	AL042615	Hs.15995	ESTs	12.35
15		Al348838	Hs.13073	ESTs	12.35
	414516	Al307802	Hs.279551	ESTs	12.34
		BE410734		gb:601301619F1 NiH_MGC_21 Homo sapiens c	12.29
		AL048542	Hs.16291	ESTs	12.28
20	401286				12.26
20		AW962845	Hs.256527		12.24
		AW512260	Hs.87767		12.24
		Al984625	Hs.9884	spindle pole body protein	12.24
	407519		Hs.25951	gb:H.sapiens mRNA HTPCRX01 for olfactory	12.23 12.22
25		AF249744 AW973653		Rho guanine nucleotide exchange factor ( hypothetical protein FLJ00052	12.20
23	405325	WW9/3003	FIS.20104	nyponencai protein Pisiooosz	12.20
		AA013140	Hs.115707	FRTs	12.18
	423066		Hs.120171		12.17
		AI623752	Hs.163603		12.16
30	443062		Hs.8963		12.15
		AA250970		Homo sapiens cDNA: FLJ23107 fis, clone L	12.14
	453542	AW836724	Hs.33190	Homo sapiens mRNA expressed only in plac	12.11
		AA864968	Hs.127699		12.10
25		AF006609	Hs.82294		12.10
35	440288		Hs.7138	cholinergic receptor, muscarinic 3	12.04
		AW024937	Hs.29410	ESTs	12.02
		AI022813	Hs.92679		
	445407	Al222658	Hs.83918	ESTs, Weakly similar to la costa [D.meta	11.95 11.94
40		Al990287	Hs.270798		11.94
40	409799		Hs.76845		11.92
		AW075485		phosphoserine amhotransferase	11.92
	443912		Hs.184780		11.92
		AA343936		gb:EST49786 Gall bladder I Homo sapiens	11.90
45		AW014795	Hs.23349	ESTs	11.90
	451533	NM_004657	Hs.26530	serum deprivation response (phosphatidyl	11.90
		AF283777	Hs.116481	CD72 antigen	11.89
		AW386461		gb:PM4-PT0019-121299-004-F02 PT0019 Homo	11.89
		AB011537	Hs.133466	slit (Drosophila) homolog 1	11.82
50		Al074413	Hs.14220		11.80
	414341	D80004	Hs.75909	KIAA0182 protein	11.80
	406538	AW450502	Hs.24218	Fev-	11.79
		BE247676	Hs.18442	ESTs E-1 enzyme	- 11.79 11.78
55		AF216751	Hs.26813	CDA14	11.76
55	416862		Hs.23575	ESTS	11.74
	425770	NM_014363		spastic ataxia of Charlevoix-Saguenay (s	11.72
		AL048842	Hs.194019	attractin	11.72
		NM 014158		HSPC067 protein	11.72
60		BE293466	Hs.20880	ESTs	11.72
		BE245374	Hs27842	hypothetical protein FLJ11210	11.72
	412922		Hs.74870	H2.0 (Drosophila)-like homeo box 1	11.72
		NM_005578		LIM domain-containing preferred transloc	11.69
65		BE548555		CGI-83 protein	11.68
U)		AF097994	ns.301528	L-kynurenine/alpha-aminoadipate aminotra	11.68
	410531 425917	AW752953	We 117107	gb:QV0-CT0224-261099-035-g02 CT0224 Homo Homo sapiens cDNA: FLJ23067 fis, clone L	11.67
		W25517 Al750878	He 87400	thrombospondin 1	11.68 11.64
	400557	M100010		ununwapallulii I	11.62
	,00001				

xog 11.60

		BE157260	Hs.79070	v-myc avian myelocytomatosis viral oncog	11.60
	419047	AW952771	Hs.90043	ESTs	11.59
		Al986160	Hs.88446	ESTs	11.59
-	400885				11.57 11.58
5	409853	AW502327		gb:UI-HF-BR0p-aka-a-07-0-UI.r1 NIH_MGC_5	11.56
		NM_016045	Un E104	TH1 drosophila homolog	11.55
		M55994		tumor necrosis factor receptor superfami	11.55
		S55736		ESTs, Weakly similar to hypothetical pro	11.54
10		AA460479	Hs.4096	KIAA0742 protein	11.53
	434228	742047	Hs 283978	ESTs: KIAA0738 gene product	11.52
	420729	AW964897 AA426080	Hs.290825	ESTs	11.52
	428328	AA426080	Hs.98489	ESTs	11.50
	433887	AW204232	Hs.279522	ESTs	11.50
15		X72755	Hs.77367		11.46
		F18572	Hs.22978		11.44
	452260	AA453208	HS.28/26	RAB9, member RAS oncogene family	11.42 11.42
	459029	AA131376	HS.2852U3	fibroblast growth factor 12 cystatin E/M	11,39
20	400207	AW975944	Hs.237396	Cysiain E/M	11.38
20		AW291876			11.37
	447861	AI434593			11.37
		R00028		gb:ye70a06.s1 Soares fetal liver spleen	11.36
		Al277652	Hs.54578	ESTs	11.31
25	401163				11.31
		L36149	Hs.248116	chemokine (C motif) XC receptor 1	11.28
		AW246803			11.28
		AL044829	Hs.29331	carnitine palmitoyltransferase I, muscle	11.27 11.26
30		NM_014253	HS.23796	odz (odd Oz/ten-m, Drosophila) homolog 1	11.24
30		AA075687 W07411	HS.14/1/6	epidermal growth factor receptor substra ESTs, Moderately similar to ALU3_HUMAN A	11.24
		H28383	118.1102.12	gb:yl52c03.r1 Soares breast 3NbHBst Homo	11.24
		AA631047	Hs 158761	Homo sapiens cDNA FLJ13054 fis, clone NT	11.23
		AA315267	Ha.23128	ESTs	11.22
35			Hs.214142	5,10-methylenetetrahydrofolate reductase	11.21
	422858	R35398		gb:yg64g10.r1 Soares infant brain 1NIB H	11.20
	415156	X84908	Hs.78060	phosphorylase kinase, beta	11.20
	446713	AV660122	Hs.282675		11.20
40	452221	C21322	Hs.11577		11.20
40		W78902	Hs.293297 Hs.127809		11.17 11.16
		Al367347 AW748078			11.16
		BE142098	H3.214410	gb:CM4-HT0137-220999-017-d11 HT0137 Homo	11.14
		AB020725	He 58009	KIAA0918 protein	11.14
45	405801	, and a second	11330000	Table to proton	11.13
		Al000341	Hs.220491		11.12
	427654	AA410183	Hs.137475		11.12
		N77624		phosphatidic acid phosphatase type 2B	11.10
<b>#</b> 0		AI567669	Hs.287316		11.10
50		AF030880		solute carrier family, member 4	11.08
		AW104057			11.07 11.07
		Y00093 W92745	Hs.51077 Hs.193324		- 11.03
		U52077	110, 100024	gb:Human mariner1 transposase gene, comp	11.02
55		AF055581	Hs.13131		11.02
		AW867079		qb:MR1-SN0033-120400-002-c10 SN0033 Homo	10.95
	401030	BE382701	Hs.25960	v-myc avian myelocytomatosis viral relat	10.95
		AW006969	Hs.6311	hypothetical protein FLJ20859	10.94
		AW591783	Hs.36131	collagen, type XIV, alpha 1 (undulin)	10.94
60		AA530994		ghrelin precursor	10.92
		AW246428	Hs.75355	ubiquitin-conjugating enzyme E2N (homolo	10.92 10.92
	400132	********	I - OTFOR	ESTs	10.92
		AA443966 NM_000328	Hs.31595	retinitis pigmentosa GTPase regulator	10.88
65		D85782	Hs.3229	cysteine dioxygenase, type I	10.88
33		Al366213		KIAA1605 protein	10.87
		AW948126		gb:RC0-MT0013-280300-031-a12 MT0013 Homo	10.85
	400615			•	10,80
	425018	BE245277	Hs.154196	E4F transcription factor 1	10.80

	456011	BE243628		gb:TCBAP1D1053 Pediatric pre-B cell acut	10.79
	455982	BE176862		gb:RC4-HT0587-170300-012-a04 HT0587 Homo	10.74
		BE218418	Hs.201802		10.73
_		AW803564	Hs.288850		10.72
5		AW377314	Hs.5364	DKFZP564l052 protein	10.70
		Al383497		ESTs, Weakly similar to ALU1_HUMAN ALU S	10.70
		R40978		ESTs, Moderately similar to ALU1_HUMAN A	10.70 10.68
	44909U	AA694070 NM_006558	Hs.13565		10.68
10		U24578		complement component 4A	10.66
10		AW863261	Hs.15036		10.64
	420090	AA220238	Hs.94986	ribonuclease P (38kD)	10.64
	451593	AF151879	Hs.26706		10.62
	438893	AF075031	Hs.29327	ESTs	10.62
15		AW080953		gb:xc28c12.x1 NCI_CGAP_Co18 Homo sapiens	10.61
		AL359652	Hs.171096	Homo sepiens EST from clone DKFZp434AD41	10.58
		AA715328	Hs.291205		10.57
		AA128423	Hs.40300	calpain 3, (p94) KIAA0128 protein; septin 2	10.57 10.56
20		D50918	HS.90998	KIAAU128 projein; septin 2	10.56
20	428522	R10184 Al142350	Hs.191987 Hs.146735	ESTs, Weakly similar to ALU1_HUMAN ALU S	10.55
		AA178955		ESTs	10.54
		AW248508			10.52
	406577				10.52
25		AK001332		hypothetical protein FLJ10470	10.51
	428966	AF059214	Hs.194687	cholesterol 25-hydroxylase	10.50
	400880				10.48
		AA894876	Hs.5687	protein phosphatase 1B (formerly 2C), ma	10.48
20		BE005346	Hs.116410		10.46
30		AA609784 Al638418		major histocompatibility complex, class	10.44 10.44
		U76421	Hs.85302		10.44
		AW500239			10,44
	419544	Alg00154		gb:QV-BT200-010499-007 BT200 Homo saplen	10.44
35	432180	Y184t8	Hs.272822	RuvB (E coll homolog)-like 1	10.44
	413822	HU8950	Hs.272044	ESTs, Weakly similar to ALU1_HUMAN ALU S	10.42
		AA788946	Hs.16869	ESTs, Moderately similar to CA1C RAT COL gamma-glutamyl hydrotase (conjugase, fol	10.41
		NM_003878	Hs.78619	gamma-glutamyt hydrotase (conjugase, fol	10.41
40		NM_003500 AW150717		acyl-Coenzyme A oxidase 2, branched chai STAT induced STAT inhibitor 3	10.38
70	415082	AA160000	Hs.137396		10.37
		AW505086		minor histocompatibility antigen HA-1	10.36
		AB011151		KIAA0579 protein	10.34
		AW067805	Hs.172665	methylenetetrahydrofolate dehydrogenase	10.34
45	424280	NM_000030	Hs.271366	alanine-glyoxylate aminotransferase homo	10.33
		T93096	Hs.17126		10.32
		NM_014324		alpha-methylacyl-CoA racemase	10.31
		AW960597	Hs.30164	ESTs, Weakly similar to ALU4_HUMAN ALU S	10.30 10.30
50		AW022715 AA172106		Rag C protein	10.30
50	406189	AA172100	115.110000	riag C protein	10.29
		AW411307	Hs.114311	CDC45 (cell division cycle 45, S.cerevis	10.28
		AA172106		Rag C protein	- 10.26
		T89832	Hs.170278		10.26
55		NM_006762	Hs.79356	Lysosomal-associated multispanning membr	10.24
			Hs.174142	colony stimulating factor 1 receptor, fo	10.24
	401384		11. 00000	and the formation of the D	10.23
		D13168 AF037062	HS.82002	endothelin receptor type B retinol dehydrogenase 5 (11-cisand 9-cis	10.22 10.21
60	428928	AI684746	Hs.119274		10.20
30		Al364997	Hs.7572	ESTS ESTS	10.20
		BE243026		KIAA0246 protein	10.19
		AA757196	Hs.121190		10.19
	403690				10.17
65		BE152393		gb:CM2-HT0323-171199-033-a08 HT0323 Homo	10.16
		AA305599		hypothetical protein PRO2013	10.16
		AW975009 Z68128	Hs.292274 Hs.3109	Rho GT Pase activating protein 4	10.16 10.16
		Al288430	Hs.64004		10.16
	-02000	7112.00400	110.04004	2010	10.14

		AW084176	Hs.223296	ESTs		10.14
		Al245701	11- 470004	gb:qk31f05.x1 NCI_CGAP_Kid3 Homo sapiens		10.13
		AA626142 Al174603		ESTs, Weakly similar to KPCE_HUMAN PROTE enclase 1, (alpha)		10.13
5		AI038989	Hs 24809	hypothetical protein FLJ10826		10.12
-		NM_006056		,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,,		10.12
		AA807346	Hs.288581	Homo sapiens cDNA FLJ14296 fis, clone PL		10.11
		AW118822	Hs.128757			10.10
		AW014605	Hs.179872	ESTs		10.10
10		H60720	Hs.81892			10.09
	442382	AI204266	Hs.179303	ESTs, Weakly similar to ALU1_HUMAN ALU S		10.05 10.04
	401202 AABBB3	RE614500	He 106823	H.sapiens gene from PAC 42616, similar t		10.04
	434467	BE552368	Hs.231853	Homo sapiens cDNA FLJ13445 fis, clone PL		10.04
15	423698	Al204266 Al433833 BE614599 BE552368 AA329796	Hs.1098	DKFZp434J1813 protein		10.02
	412707	AW206373	Hs.16443	Homo sapiens cDNA: FLJ21721 fis, clone C		10.00
		X58528		ATP-binding cassette, sub-family D (ALD)		10.00
		NM_016098		HSPC040 protein		10.00
20	423554	M90516 Al922988	Hs.1674 Hs.172510	glutamine-fructose-6-phosphate transamin		10.00
20	402009	Al922988 AW137442 AA418280 BE501815	Hs.138965			10.00
	427978	A4418280	Hs 180040	Home sapiens cDNA: FLJ22439 fis, clone H		10.00
	457803	BE501815	Hs.198011			9.99
	428279	AA425310	Hs.155765			9.98
25	444412	Al147652		Homo saplens clone HH409 unknown mRNA		9.98
		N72394	Hs.44862			9.96
		M62505	Hs.2161	complement component 5 receptor 1 (C5a I		9.96
	445424	AB028945 AW009605	HS. 12090	cortactin SH3 domain-binding protein		9.96 9.96
30	443070	AW474513	He 224397	ESTs, Weakly similar to B48013 proline-r		9.94
-	414709	AA704703	Hs.77031	Sp2 transcription factor		9.94
	434506	TF0538		ob vh85g12 s1 Stratagene ovary (937217)		9.94
	427630	BE276115	Hs.144980	ESTs, Weakly similar to CA13_HUMAN COLLA		9.93
25	416111	AA033813	Hs.79018	ESTs, Weakly similar to CA13_HUMAN COLLA chromatin assembly factor 1, subunit A ( homeo box A9		9.92
35	423349	AF010258	Hs.127428	homeo box A9		9.92
	424308	AW975531 AW192307	HS.154443	minichromosome maintenance deficient (S. dollchyl-P-Glo:Man9GloNAc2-PP-dollchylgi		9.92
	410014 417088	AA481003	Hs.97128			9.90
	425174	AA481003 D87450	Hs.154978	KIAA0261 protein		9.90
40		AW976507	Hs.293515			9.90
	421984	AW972187	Hs.110443	hypothetical protein FLJ22215		9.89
		NM_005291		G protein-coupled receptor 17		9.88
		AI097570	Hs.71222	ESTs		9.87
45		AW801383 AI278802	Hs.25661	H.sapiens mRNA for ribosomal protein L18 ESTs		9.86 9.85
43		AW117416	Hs.245484			9.85
		AL043002		ESTs. Moderately similar to unnamed prot		9.84
	449824	AI962552	Hs.226765			9.84
	452744	Al267652	Hs.30504	Homo sapiens mRNA; cDNA DKFZp434E082 (fr		9.82
50	431066	AF026273	Hs.249175	Interleukin-1 receptor-associated kinase		9.82
	426457	AW894667		chimerin (chimaerin) 1		9.80
		Al792888	Hs.145489	gb:Homo sapiens mRNA; cDNA DKFZp566E1346		9.80 9.75
		AL050072 D13635	He 155287	KIAA0010 gene product	•	9.74
55		N67819	Hs.43687	ESTs		9.74
		Al590401	Hs.21213	ESTs		9.73
		Al381659	Hs.267086			9.72
		AW748265	Hs.5741	flavohemoprotein b5+b5R		9.72
60		AL120659	Hs.6111	KIAA0307 gene product		9.72
60		AA807228 AA311152	Hs.225161	ESTS; Weekly similar to KIAA0226 [H.sapl		9.72 9.72
		Al289619	Hs.13040			9.72
	436209	AK001451	Hs.265561	CD2-associated protein		9.70
	446896	T15767	Hs.22452	Homo sapiens cDNA: FLJ21084 fis, clone C		9.70
65	412667	AW977540	Hs.269254			9.70
		S67580	Hs.1645	cytochrome P450, subfamily IVA, polypept		9.67
	440757	AW118645				9.67
		Al393657 AF061871	Hs.159750 Hs 101302	collagen, type XII, alpha 1		9.66 9.66
	72 7044	A 00 10/ I	1201002	countries the suit advise 1		0.00

		BE466863	Hs.280099	ESTs	9.68
		R91679	Hs.124981		9.66
		X02422	Hs.181125	immunoglobulin lambda locus	9.65
	441530	Al248301	Hs.127112	ESTs	9.65
5		D53304	Hs.65394		9.65
	421470	R27496	Hs.1378	annexin A3	9.64
		C05569	Hs.243122	hypothetical protein FLJ13057 similar to	9.64
	429324	AA488101		inactivation escape 1	9.62
	450244	AA007534	Hs.125062	ESTs	9.62
10	407660	AW063190	Hs.279101	ESTs	9.61
	406554				9.60
	426404	AA377607	Hs.273138	ESTs	9.58
	447045	AW392394	Hs.278569	KIAA0064 gene product	9.58
	449894	AK001578	Hs.24129	hypothetical protein FLJ10716	9.58
15	448376	Al494332	Hs.196963	ESTs	9.58
		AL117474		Homo sapiens mRNA; cDNA DKFZp727C191 (fr	9.56
	446572	AV659151	Hs.282961	ESTs	9.58
		BE242623		manic fringe (Drosophila) homolog	9.55
		AP000692		chromosome 21 open reading frame 5	9.54
20	414697	BE266134	Hs.76927	translocase of outer mitochondrial membr	9.54
	410848	AW807057		gb:MR4-ST0062-031199-018-b03 ST0062 Homo	9.52
		NM_005574		LIM domain only 2 (rhombotin-like 1)	9.52
	427308		Hs.174905	KIAA0033 protein	9.52
		NM_004573	Hs.994	phospholipase C, beta 2	9.51
25		AW295389	Hs.119768		9.51
		AA742181	Hs.75912	Homo sapiens cDNA: FLJ22199 fis, clone H	9.50
		D28459	Hs.80612	ubiquitin-conjugating enzyme E2A (RAD6 h	9.50
		AA094538	Hs.6588	ESTs	9.50
		AA833902	Hs.270745		9.48
30		R07114	Hs.271224	ESTs	9.48
		AJ132085		gb:Homo sapiens mRNA for axonemal dynein	9.44
		AW137726		ESTs, Moderately similar to laminin alph	9.44
		AW450584	Hs.192131	ESTs, Weakly similar to RIBB [H.saplens]	9.43
~~	404741				9.43
35		NM_005428	Hs.116237	vav 1 oncogene	9.43
	403708				9.42
		AW847814	Hs.289005	Homo sapiens cDNA: FLJ21532 fis, clone C	9.42
	417380			gb:EST04698 Fetal brain, Stratagene (cat	9.42
40		AA354690	Hs.144967		9.42
40		AA004410		acyl-Coenzyme A oxidase 1, palmitoyi	9.42
		AU076606	HS.30054	coagulation factor V (proaccelerin, labi	9.42
		AW893569		gb:RC0-NN0021-040400-021-c10 NN0021 Homo	9.41
		AA361623		Homo sapiens cDNA FLJ13900 fis, clone TH	9.41
45	408101	AW968504		CDC2-related protein kinase 7	9.40 9.40
45		AA360328	Hs.865	RAP1A, member of RAS oncogene family mutS (E. coll) homolog 2 (colon cancer,	9.40
		U04045 BE262745	HS./8934		9.40
		AI689987	11- 477000	gb:601153869F1 NIH_MGC_19 Homo saplens c	9.39
		BE514362	HS.177009	ESTs, Weakly similar to RMS1_HUMAN REGUL	9.39
50	402835	DE014302	NS-200422	FK506-binding protein 3 (25kD)	9.36
50	404632				9.36
		H95741	He 17014	Homo sapiens cDNA: FLJ22801 fls, clone K	9.37
		AW903533	115.17014	gb:CM1-NN1031-060400-178-d05 NN1031 Homo	- 9.37
		AW90333 AI095087	He 152200	ESTs, Moderately similar to ALU5_HUMAN A	9.36
55		Al420611	Hs.127832		9.36
55		BE258532		CTP synthase	9.34
	120227	AA283G81		prostaglandin E receptor 4 (subtype EP4)	9.33
	407081	AA283981 X97748	113.1002-43	ghth.sapiens PTX3 gene promotor region.	9.33
	416967	BE616731	Hs.80645		9.33
60		AW875443		secreted modular calcium-binding protein	9.33
		AA693960	Hs.103158		9.33
		BE513731		Human DNA sequence from clone 967N21 on	9.32
		AA033699		ESTs, Moderately, similar to MASP-2 [H.sa	9.32
		NM_007274	Hs.8679	cytosolic acyl coenzyme A thicester hydr	9.32
65	452859	Al300555		Homo sapiens cDNA: FLJ23591 fis, clone L	9.32
-	403237				9.32
		AMMOREON	Un 220012	ESTs, Weakly similar to CALM_HUMAN CALMO	9.31
	415000				
		AW976410	Hs.289069	Homo saplens cDNA: FLJ21016 fis, clone C	9.30
	417951		Hs.289069 Hs.6975	Homo saplens cDNA: FLJ21016 fis, clone C PRO1073 protein	9.30 9.30

414726 BE466863 Hs 280099 ESTs

		AW167128	Hs.231934	ESTs	9.30
	405125				9.30
		AW499566		gb:UI-HF-BR0p-aji-h-03-0-UI.r1 NIH_MGC_5	9.28
5		Al191811	Hs.54629	ESTs	9.28 9.27
,	442271	AF000652	Ho 50244	syndecan binding protein (syntenin) gene for serine/threorine protein kinase	9.26
	448899	AW013907	He 224278	gene for serine/threorine protein kinase ESTs, Moderately similar to predicted us solute carrier family 23 (nucleobase tra	9.26
	/17791	AF184142	He 82042	solute carrier family 23 (nucleobase tra	9.25
				KIAA0053 gene product	9,25
10	414140	AA201270	Un 22217	ECTA	0.24
	435980	AF274571	Hs.129142	ESTs; Weakly similar to DEOXYRIBONUCLEAS	9.24
	458530	BE395035	Hs.199869	ESTs; Weakly similar to DEOXYRIBONUCLEAS ESTs, Weakly similar to KIAA0874 protein	9.24
	402585				9.24
	420819	AA280700		gb:zs95h11.s1 NCI_CGAP_GCB1 Homo saplens	
15	444755	AA431791 U42349	Hs.183001	ESTS	9.22
	411630	U42349	Hs.71119	Putative prostate cancer tumor suppresso	9.22
	421246	AW562962	MS.300901	ESTs, Highly similar to AF151805 1 CGI-4	9.20
	421924	AL030165	HS.103000	coronin, actin-binding protein, 1A thyroid hormone receptor interactor 7 ESTS	9.19 9.18
20	424267	VI308280	He 116243	FSTe	9.17
20	409213	U61412	Hs.51133	PTK6 protein tyrosine kinase 6 leukemia inhibitory factor (cholinergio	9.17
		H55709	Hs.2250	leukemia inhibitory factor (cholineralo	9.18
	451736	AW080356	Hs.293684	ESTs, Weakly similar to alternatively sp	9.15
	413627	BE182082 AA528402	Hs.246973	ESTs	9.14
25	416134	AA528402	Hs.74861	activated RNA polymerase II transcriptio	9.14
	449251	AW151660	Hs.31444		9.14
	452813	U54727 Al911527	Hs.191445	ESTs	9.14
	443622	AI911527	Hs.11805	ESTS	9.14
30	413260	BE075281		gb:PM1-BT0585-290200-005-d07 BT0585 Homo	9.12
30	413450	Z99/16	H8./53/2	N-acetylgalactosaminicase, alpha-	9.12 9.12
	490540	AA010001	Hs.237000	ECTA	9.12
	426251	M24283	He 168383	SD18 Sp.PMI-BT0SSS-290200-005-d07 BT0S85 Homo N-acetylgaladosaminidase, alpha- EST3 EST3 Intercellular adhesion molecule 1 (CD54) ubiquinol-cytochrome o reductase hinge p EST3	9.11
	410290	AA402307	Hs.73818	ubiquinol-cytochrome c reductase hinge p	9.10
35	437398	AA913736	Hs.126715	ESTs	9.10
				Stezu-related senne/in/eonine kinase	9.10
	439699	AF086534 C19035	Hs.187561	ESTs, Moderately similar to ALU1_HUMAN A	9.10
	430799	C19035 M88700	Hs.164259		9.09
40				dopa decarboxylase (aromatic L-amino aci	9.08
40	453942	AW190920	Hs.19928	ESIS	9.08
	420844	T68073 Al624436 BE328153	M8.108028	serine (or cysteine) proteinase inhibito	9.08 9.07
	454000	MI024430	He 240087	ESIS ESTo	9.06
	436490	B71543	Hs 18713	EST'S EST'S, Weakly similar to ALUT_HUMAN ALU S hypothetical protein FL112999 Homo saptens cDNA FL113999 fs, clone NT Homo saptens mRNA; cDNA DKFZp654D1164 (f EST'S EST SA ORDER)	9.05
45	409192	AA065131	Hs.233439	ESTs. Weakly similar to ALU7 HUMAN ALU S	9.05
	446223	BE300091	Hs.119699	hypothetical protein FLJ12959	9.04
	447247	AW369351	Hs.287955	Homo saptens cDNA FLJ13090 fis, clone NT	9.04
	450094	Al174947	Hs.295789	Homo saplens mRNA; cDNA DKFZp664D1164 (f	9.04
~~	432012	AW301344	Hs.195969	ESTs	9.04
50	422520	AU076730 BE396750 M81590	Hs.117977	MIDSHI & (OUTUND)	9.02
	418650	BE396750	HS.86978	prolyl endopeptidase	9.02
	423008	AA326108	MS.123016	5-hydroxytryptamine (serotonin) receptor	- 9.02
	440000	BE622585	H8.00001	ESTS	9.02
55	431574	AW572650	Hs 261373	adenosine A2h recentor oseudogene	9.01
55	443453	R99878	Hs 269882	adenosine A2b receptor pseudogene ESTs triggering receptor expressed on myeloid	9.01
	435472	AW972330	Hs.283022	tripgering receptor expressed on myeloid	9.01
	420337	AW295840	Hs.14555	Homo saniens cDNA: FLJ21513 fis. clone C	9.00
	449810	AB038681	Hs.23994	activin A receptor, type IIB rlbosomal protein L4	9.00
60	406780	AA902386	Hs.286	ribosomal protein L4	8.99
					8.99
	421326	AF051428	Hs.103504	estrogen receptor 2 (ER beta)	8.97
	425491	A4883316	MS.255221	ESIS FOT-	8.96
65	420516	AI051219	∏8.2355/ He 1/331∈	ESTe	8.96 8.96
0.5	443247	RE61/397	He 47378	FSTs	8.96
	456623	AW341130 AF051428 AA883316 BE000707 AI051313 BE614387 AI084125 L06239	Hs.108106	transcription factor	8.95
	438707	L06239	Hs.5326	porcupine	8.95
	402240			•	8.95

	444152	Al125694	Hs.149305	Homo sapiens cDNA FLJ14264 lis, clone PL	8,95
	409842	AW501756		gb:UI-HF-BR0p-ajm-c-09-0-Ul.r1 NIH_MGC_5	8.94
		W78765	Hs.73580	ESTs	8.94
_	456697	A1908006		ferritin, light polypeptide	8.94
5		AF226053		HSKM-B protein	8.92
		AL120344	Hs.75074	mitogen-activated protein kinase-activat	8.92
		Al287817	Hs.129636		8.92
		AA002064	Hs.18920	ESTs	8.91
10	411486			eukaryotic translation elongation factor	8.90
10		BE566249	Hs.20999		8.90
		AA257161	Hs.8658	hypothetical protein DKFZp434E0321	8.89 8.89
		NM_007019	HS.93002	ubiquitin carrier protein E2-C	8.89
		AW849292	Hs.290259	gb:lL3-CT0215-020300-090-E06 CT0215 Homo	8.89
15	431154	AW971228	Hs.77631	glycine cleavage system protein H (amino	8.88
13	418036		Hs.83337	latent transforming growth factor beta b	8.87
	406422	231910	118.00001	rateur nausionning grown racio: beta b	8.87
		NM_016102	He 121748	ring finger protein 16	8.87
		D50030	Hs.104	HGF activator	8.86
20	418203		Hs.83758		8.86
20	418613	AA744529	Hs.86575		8.85
		H66566	Hs.271711		8.85
		AA076049	Hs.274415	Homo saplens cDNA FLJ10229 fis, clone HE	8.84
		AI952797		Homo saplens cDNA: FLJ21559 fis, clone C	8.83
25		T89839	Hs.119471	ESTs	8.83
	425894	U51333		hexokinase 3 (white cell)	8.82
		AL041465		ESTs, Moderately similar to ALU2_HUMAN A	8.82
		Al683487		Homo sapiens cDNA FLJ11441 fis, clone HE	8.82
	413413	D82520	Hs.301834	Homo sapiens cDNA FLJ10952 fis, clone PL	8.82
30	428907	AA435997	Hs.104930		8.82
		R40611	Hs.137565		8.81
		N34145	Hs.250614		8.80
		AW043637	Hs.21766	ESTs	8.80
~~		Al952677		Homo sapiens mRNA; cDNA DKFZp434P228 (fr	8.80
35		AW292577	Hs.94445	ESTs	8.80
		BE063380		gb:PM0-BT0275-291099-002-g10 BT0275 Homo	8.80 8.78
	403259	************		-1-11 0 070010 000100 001 B11 070010 H	8.78
		AW850473 BE264645	11- 000000	gb:lL3-CT0219-280100-061-B11 CT0219 Homo Homo sapiens cDNA: FLJ21918 fis, clone H	8.77
40				peroxiredoxin 3	8.76
40	401744	AA331861	H5./ 0404	peroxieuoxii S	8.76
		AL137477	He 155012	cadherin-like 24	8.76
		Al382555		bromodomain-containing 1	8.75
		NM_001429	Hs.297722	Human DNA sequence from clone RP1+85F18	8.75
45		NM_007240	Hs.44229	dual specificity phosphatase 12	8.74
		AB020316	Hs.134015	uronyl 2-sulfotransferase	8.74
		A1566086	Hs.153716	Homo sapiens mRNA for Hmob33 protein, 3'	8.74
		AA449506	Hs.179765	Homo saplens mRNA; cDNA DKFZp586H1921 (f	8.73
		AA551010	Hs.216640		8.72
50		AL137527	Hs.22703	Homo sapiens mRNA; cDNA DKFZp434P1018 (f	8.72
	429455	Al472111	Hs.292507		8.71
		AV/385597		ESTs, Weakly similar to B34087 hypotheti	8.71
	441748	H59955	Hs.127829		8.70
		AL033527		v-myc avian myelocytomatosis viral oncog	8.70
55		D87470		KIAA0280 protein	8.70
		W31254	Hs.7045	GL004 protein	8.70
		AA609019	Hs.159343		8.70
		Z97989		FYN oncogene related to SRC, FGR, YES	8.69
60		AA317036	Hs.41989		8.67
60		Al225235		Homo sapiens cDNA: FLJ23231 fis, clone C	8.67 8.66
		AA811813	Hs.119421		8.66
		AA256756	Hs.31178	ESTs granzyme K (serine protease, granzyme 3;	8.66
		NM_002104	Hs.3066	Homo sapiens mRNA for FLJ00020 protein,	8.65
65		BE244076 BE246449	Hs.2157		8.64
03		W68180	He 250055	Homo sapiens cDNA FLJ12507 fis, clone NT	8.64
		AJ001443	He 195014	splicing factor 3b, subunit 3, 130kD	8.64
		NM_006895	Hs.81182		8.64
				xenotropic and polytropic retrovirus rec	8.63
	-10/2/1				0.00
				106	

	422631	BE218919	Hs.118793	hypothetical protein FLJ10588	8.63
	410679	AW795196	Hs.215857	ring finger protein 14	8.83
	431585	BE242803	Hs.262823	hypothetical protein FLJ10326	8.62
	401851				8.62
5	401866				8.62
		AW996872		a disintegrin and metalloproteinase doma	8.62
		AA251594		PIBF1 gene product	8.62
		AW408530		ClpX (caseinolytic protease X, E. coli)	8.62
10		BE550182		RalGEF-like protein 3, mouse homolog	8.62
10		Al831594	Hs.68647		8.62 8.60
		AW749617	Hs.82302	gb:RC3-BT0502-130100-012-g07 BT0502 Homo	8.60
		Al787758	Hs.47939	ESIS For-	8.60
	429328	AA829402		Homo sacions cDNA FLJ13741 fis, clone PL	8.60
15	400004	Al972094 Al692181	No 40100	KIAA1634 protein	8.60
13	400007	AF009746 X54136	He 0/306	ATP-binding cassette, sub-family D (ALD)	8.60
	4250027	Y54198	He 181125	immunoglobulin lambda locus	8.60
	430000	U91939	He 248123	G protein-coupled receptor 25	8.60
	405074	001000	TICLE TO LEC	a protest company and	8.59
20		Al479773	Hs.181679	ESTs	8.59
20		BE328882		ESTs, Moderately similar to U119_HUMAN U	8.58
	411079	AA091228		gb:ochn2152.seq.F Human fetal heart, Lam	8.57
	418452	BE379749	Hs.85201	C-type (calcium dependent, carbohydrate-	8.56
		AL009637		neutrophil cytosolic factor 4 (40kD)	8.56
25		AW947164	Hs.195641		8.56
		AW204272	Hs.199371		8.55
		H55883		gb:yq94h03.r1 Soares fetal liver spleen	8.54
		BE007663	Hs.13503	inactivation escape 2	8.54
20	405876				8.54 8.54
30		D20569	HS.169407	SAC2 (suppressor of actin mutations 2, y	8.54
		Al738616 AF193612	HS.//348	hydroxyprostaglandin dehydrogenase 15-(N lunatic fringe (Drosophila) homolog	8.54
		AP193612 AW082633	Hs.212715		8.54
		AA446183	Hs.91885		8.53
35		AI955765	Hs.146907		8.52
55		M31899	Hs.77929		8.51
	405552	MOTODO	110H TOLK	and the state of t	8.51
		AW971155	Hs 293902	ESTs, Wealdy similar to prolyl 4-hydroxy	8.50
		AA426117	Hs.14373		8.50
40		R68857	Hs.265499	ESTs	8.50
		Al765890	Hs.16341	ESTs; Moderately similar to !!!! ALU SUB	8.50
		AV659397	Hs.282948		8.50
		AW891873		gb:CM3-NT0090-040500-173-b02 NT0090 Homo	8.50
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45		AA279098	Hs.187636		8.48 8.48
		AW137635	Hs.44238		8.48
		AL043983	Hs.125063	Homo sapiens cDNA FLJ13825 fis, clone TH	8,48
		Al907957	HS.28/622	Homo sapiens cDNA FLJ14082 fis, clone HE	8.48
50		AA229126	Ms. 122047	N-myristoytransferase 2 CD36 antigen (collagen type I receptor,	8.47
50		AA593731 AL038704	Ho 150007	ESTs, Weakly similar to ALU1_HUMAN ALU S	8.46
		AL030704 AL080276	He 269662	regulator of G-protein signalling 17	8.46
	400268	ALUGUZIO	110.200002	regulator or a protein agreeing 17	- 8.46
		AW891965	Hs 289109	dimethylarginine dimethylaminohydrolase	8.45
55		NM_014737	Hs.80905	Res association (RalGDS/AF-6) domain fam	8.44
-		AW057782	Hs.293053	ESTs	8.44
	425171	AW732240	Hs.300615		8.44
		Al814302		gb:wj71c12.x1 NCI_CGAP_Lu19 Homo saplens	8.42
	406006				8.42
60	412643	AW971239	Hs.293962		8.42
		AB014540	Hs.153026	SWAP-70 protein	8.42
		AW136083		ESTs, Weakly similar to \$59501 interfero	8.42
		AI458653	Hs.201881	ESTS	8.41
		AA358015		gb:EST66864 Fetal lung III Homo sapiens	8.40
65		AW978439	Hs.69504		8.40 8.40
		AA013051	Hs.91417	gb:EST384925 MAGE resequences, MAGL Homo	8.40
		AW972830 AA305688	No 007005	UDP-Gal:betaGlcNAc beta 1,3-galactosyltr	8.40
		AI521310	He 283365	ESTs, Weakly similar to ALU5_HUMAN ALU S	8.40
	40000	1010			

	447685	AL122043	Hs.19221		8.40
	459119	AW844498	Hs.289052	Homo saplens LENG8 mRNA, variant C, part	8.38
	400817				8.37
	425265	BE245297		gb:TCBAP1E2482 Pediatric pre-B cell acut	8.37
5	409385	AA071267		gb:zm61g01.r1 Stratagene fibroblast (937	8.36
	439121	BE047779	Hs.44701	ESTs	8.36
		X04430		interieukin 6 (interferon, beta 2)	8.36
		AW182309	Hs.249963	ESTs, Highly similar to dJ1170K4.4 [H.sa	8.35
	403976				8.34
10		AA379036		gb:EST91809 Synovial sarcoma Homo sapien	8.33
		AW188551			8.33
		AW997704		Hamo sapiens cDNA FLJ13536 fis, clone PL	8.32
		AF119847		Homo sapiens PRO1550 mRNA, partial cds	8.32
1.5		AW937670	Hs.254379		8.32
15		NM_015698	Hs.100391	T54 protein	8.30
		T70298		gb:yd26g02.s1 Soares fetal liver spleen	8.30 8.30
		AF283776	Hs.80285		8.30
		AF084866	11-00500	gb:Homo sapiens envelope protein RIC-3 ( ESTs	8.29
20		Al732694	Hs.98520 Hs.199028		8.29
20		AW194962 BE266695	HS. 188028	gb:601190242F1 NIH MGC 7 Homo sapiens cD	8.29
	404946	BE200093		go.oorroozezi i Niir_wao_r nano sapiene co	8.28
		AF054839	U- 100E40	tetraspan 2	8.28
		AA037675	Hs.152675		8.28
25		AA744488		ESTs, Moderately similar to ALU1_HUMAN A	8.28
23		AU076484	Hs.9963	TYRO protein tyrosine kinase binding pro	8.27
		AF106069	He 23168	ubiquitin specific protease 15	8.26
		AA151730		ESTs, Weakly similar to similar to C.ele	8.26
		AB007918		KIAA0449 protein	8.25
30		AA974411	Hs.18672		8.25
		AW958264	Hs.103832	ESTs, Weakly similar to TRHY_HUMAN TRICH	8.24
		Al963740	Hs.46826	ESTs	8.24
	427359	AW020782	Hs.79881	Homo sapiens cDNA: FLJ23006 fis, clone L	8.24
	424534	D87682	Hs.150275	KiAA0241 protein	8.24
35		U63830		TRAF family member-associated NFKB activ	8.24
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40		BE222450	Hs.266390		8.21
40		AA330214		gb:EST33935 Embryo, 12 week II Homo sapi	8.21
		AA888624	Hs.19121		8.20
		AA835868		Homo sapiens cDNA: FLJ20935 fis, clone A	8.20 8.20
		R40739	Hs.21326		8.20
45		W25760	Hs.77631	glycine cleavage system protein H (amino minichromosome maintenance deficient (S.	8.20
43		AU077143 AV654978		cystathionase (cystathionine gamma-lyase	8.19
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		NM 015896		BLu protein	8.18
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50		M21305		Human alpha satellite and satellite 3 ju	8.18
50	400304	U42349	Hs.71119		8.18
	423397	NM 001838	Hs.1652	chemokine (C-C motif) receptor 7	8.18
		AL133017	Hs.2210	thyroid hormone receptor interactor 3	- 8.17
	401519				8.17
55		H65423	Hs.17631	Homo saplens cDNA FLJ20118 fis, clone CO	8.16
	424704	Al263293	Hs.152096	cytochrome P450, subfamily IIJ (arachido	8.16
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		AA278536	Hs.23262		8.14
60		AI139058	Hs.23296	ESTs	8.14
		Al018408	Hs.131284		8.14
	421129	BE439899	Hs.89271	ESTs	8.14

TABLE 9A shows the accession numbers for those primekeys lacking a unigeneID in Table 9. For each probeset we have listed the gene cluster number from which the oligonucleotides were designed. Gene clusters were compiled using sequences derived from Genbank ESTs and mRNAs. These sequences were clustered based on sequence similarity using Clustering and Alignment Tools (Double Twist, Oakland California). The Genbank accession numbers

for sequences comprising each cluster are listed in the "Accession" column.

5

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        CAT number:
                           Genbank accession numbers
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	450783	84655_1	BE266695 BE265474 N53200 BE267333
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	452549	921802 1	Al907039 Al907061
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	457595	3842251	AA584854
50	457751	399422_1	AI908236 AA663731
		883688 1	Al814302 Al814428
		889426_1	W07608 AIS22066
		918957_1	AI903354 AI903489 AI903488
		921149_1	BE063380 BE063346 Al906097
55		9452401	AB40425
	100207		

TABLE 9B shows the genomic positioning for those primekeys lacking unigene ID's and accession numbers in Table 9. For each predicted exon, we have listed the genomic sequence source used for prediction. Nucleotide locations of each predicted exon are also listed.

Unique number corresponding to an Eos probeset Pkey: Sequence source. The 7 digit numbers in this column are Genbank Identifier (GI) numbers. "Dunham I. et al." refers to the Ref: publication entitled "The DNA sequence of human chromosome 22." Dunham I. et al., Nature (1999) 402:489-495. Indicates DNA strand from which exons were predicted. Strand: 10 indicates nucleotide positions of predicted exons. Nt\_position: Pkey Ref Strand Mt position 15 400452 8113550 Minus 90308-90505 400557 9801261 208453-208528,209633-209813 Plus 400615 9908994 Plus 118036-118166.118881-118807 400802 8567867 Minus 174571-174856 170793-170948 400817 8569994 Plus 20 400880 9931121 29235-29336.36363-36580 Plus 400885 9958187 Minus Minus 52033-52158.53958-54120.54957-55052.55420-55480.56452-56666.57221-57718 400926 7651921 400952 7658481 Plus 192667-192826.194387-194876 400991 8096825 Plus 159197-159320 25 401044 8117619 Plus 73501-73674 Minus 124181-124391 401124 8570298 5302-5545 401163 6981820 Pine 138534-138629.139234-139294,140121-140335,142033-142479 401201 9743387 Minus 401286 9801342 Minus 147036-147318 30 Minus 58360-58545 401384 6850939 401468 6433326 Plus 13056-13482 401515 7630851 Plus 29929-30126 157315-157950 401519 6649315 Plus Plus 128526-128704.130755-130860 401672 9838136 35 401744 2576349 Plus 14595-14751 146443-146664.147794-147971.148351-148480.148980-149111.149801-149949 401851 7770425 Minus Plus 73126-73623 401866 8018106 402240 7690131 Plus 104382-104527,106136-106372 402359 9211204 Minus 40403-41961 Minus 174893-175050 183210-183435 402585 9908890 402788 9796102 Plus 98273-101430 53242-53432 402802 3287156 Minus 25026-25091,25844-25920 402812 6010110 Plus 402828 8918414 Plus 69071-69642 45 402835 9187337 Plus 26961-27101 402838 9369121 Minus 32589-32735,35478-35666 Minus 78355-76479 402842 9369121 85537-85671.86379-86469 402895 9967547 Plus 402984 9581599 Minus 46824-46784 50 Minus 92349-92572,92958-93084,93579-93712,93949-94072,94591-94748,95214-95337 403137 9211494 403237 7637807 Plus 7271-7527 403259 7770585 Plus 4693-4857 403883 7331517 Plus 217175-217446 Minus 78627-79583 403690 7387384 55 134394-134812 403708 5705981 Minus Plus 19197-19502 403838 4178355 22733-23007 403851 7708872 Plus 403976 7657840 Plus 24755-24969 404407 7329316 Minus 48154-48499 404426 7407959 Dine 77842-77954 404632 9796668 Plus 45098-45229 404741 8574139 Plus 143025-143467 82849-83627 404756 7706327 Plus Plus 134445-134750 A010A6 7382180 65 44340-44559,44790-45059 405074 7770440 Plus

405125 8247873 Plus

405172 9966752 Plus

137113-137814

153027-153262

	405236 7249076	Minus	151699-151915
	405325 6094861	Minus	25818-26380
	405411 3451356	Minus	17503-17778,18021-18290
	405495 8050952	Minus	72182-72373
5	405552 1552506	Plus	45199-45647
	405601 5815493	Minus	147835-147935,149220-149299
	405685 4508129	Minus	37956-38097
	405777 7263187	Minus	104773-105051
	405856 7653009	Plus	101777-102043
10	405876 6758747	Plus	39694-40031
	405932 7767812	Minus	123525-123713
	405934 6758795	Plus	159913-160605
	406006 8247801	Minus	42640-42776
	406134 9163473	Plus	153291-153452
15	406189 7289992	Minus	22007-22234
	406422 9256411	Plus	163003-163311
	406516 7711422	Minus	128375-128449.128560-128784
	406538 7711478	Plus	35196-35367,38229-38476,40080-40216,43522-43840
	406554 7711566	Plus	106956-107121
20	406577 7711730	Pius	11377-11509

TABLE 10: shows genes, including expression sequence tags differentially expressed in taxol resistant prostate tumor xenografts as compared to taxol sensitive prostate tumor xenografts. The genes are indicated as either being upregulated or downregulated during the induction of taxol resistance in sequential passages of the grafts.

10 15	Pkey: ExAcon: Unigene Unigene Eos: F00-F14	iD: Title:	Unique Eos probesel identifier number Exemplar Accession number, Genkank accession number Unique number Unique neges title Iniquen eges title Iniquen eges title Iniquente passe title Iniquente passe title Iniquente passe																	
	Pkey	ExAcon	UnigenelD	UnigenTitle	Eos	Resp	.F00	F00	F02	F02	F05	F05	F07	F09	F10	F11	F13	F14		
20	117921	N51002	Hs.47170	Liprin A2	PM2	RUP	1	9	8	9	32	20	34	122	105	82	71	111		
		T17185	Hs.4299	ESTs		1 down	290		267	335	270	284	150	157	83	89	49	75		
		Al167942	Hs.61635	STEAP		down			103	71	34	67	33	14	2	1	1	1		
		N95796	Hs.179809	FSTs	PAB	down	765	841	757	909	742	704	478	428	253	175	228	238		
		N31952	Hs.167531			down			147	141	123	129	73	65	55	48	54	84		
25		HG2841-HT29		Hs.75442	Albut	nin. A	PMO:	i down	666	605	504	728	357	445	602	187	117	127	117	113
		HG2841-HT29		Hs.75442				down		653	486	688	368	386	606	175	101	95	115	97
		U09579	Hs.252437			3down		94	143	190	105	107	88	40	34	31	46	22		
		U22961	Hs.75442	aibumin		down		424	323	518	252	296	467	188	169	143	165	145		
		AA075779	•	mitochondr				190	606	230	378	106	218	88	69	192	69	99		
30		AA599690	Hs.15725	SBBI48		down			115	188	132	111	66	71	49	70	38	50		
50		AA062746	-	ESTs		7down			252	13	22	43	193	10	10	104	21	18		
		AA065143	-	solute car	PMO	3down	27	54	178	73	108	37	53	24	14	53	15	34		
		AA115963				down		893	1292	656	869	389	1	74	118	662	359	409		
		AA126313	Hs.107476	ATP syntha	PM1	down	10	19	185	25	60	1	32	3	7	14	1	1		
35		H89355	Hs.6598	adrenergic					237	239	231	220	119	145	93	64	56	124		
55		AA283804	Hs.193552			down			282	271	340	334	115	238	100	196	83	207		
		AA430124	Hs.234607	MDM2	PM13	down	49	93	94	154	132	91	23	54	23	76	14	41		
		AA281591	Hs.16193	ESTs	PM1	down	80	157	58	141	159	127	39	83	35	37	16	46		
		Y00705	Hs.181286	serine pro	PMI	down	146	217	214	150	108	128	177	85	54	63	66	56		
40		AA490775	Hs.5920	N-acetylma				150	132	178	126	139	53	94	48	67	41	80		
		AA032221	Hs.61635	STEAP		down			203	215	205	180	132	65	68	50	48	63		
		AA283085	Hs.64065	ESTs		Bdown		148	161	150	92	108	42	99	42	65	29	126		
		D62633	Hs.8238	ESTs		dawn			194	212	231	189	89	123	107	95	68	91		
		M23263	Hs.99915	androgen r		Odown		167	99	178	132	101	23	71	28	122	14	44		
45																-				

TABLE 11: shows genes, including expression sequence tags that are up-regulated in prostate tumor tissue compared to normal prostate tissue as analyzed using Affymetrix/Eos Hu01 GeneChip array. Shown are the ratios of "average" normal prostate to "average" prostate cancer tissues.

5

	Pkey:		set identifier number		
	ExAcon:		ion number, Genbani	k accession number	
	UnigeneiD:	Unigene number			
	Unigene Title: B1:			a complete transferre	
	- Н1:	Background subt	racted normal prostat	e : prostate tumor tissue	
	Pkey	ExAcon	UnigenelD	Unigene Title	R1
	101336	L49169	Hs.75678	FBJ murine osteosarcoma viral oncogene homolog B	0.012
10	130642	M63438	Hs.156110	Immunoglobulin kappa variable 1D-8	0.015
	133512	X01677	Hs.195188	glyceraldehyde-3-phosphate dehydrogenase	0.017
	133436	H44631	Hs.737	immediate early protein	0.017
	129292	X13810	Hs,1101	POU domain; class 2; transcription factor 2	0.019
	100610	HG2566-HT4792		Microtubule-Associated Protein Tau, Alt. Splice 3, Exon 8	0.02
15	133448	M34516	Hs.170116	immunoglobulin lambda-like polypeptide 3	0.021
	125193	W67577	Hs.84298	CD74 antigen (invariant polypeptide of major histocompatibility	
				complex class II antigen-associated)	0.022
	133456	T49257	Hs.183704	ubiquitin C	0.022
	134546	AA459310	Hs.8518	Homo sapiens mRNA; cDNA DKFZp586L1722 (from clone	
20				DKFZ0586L1722)	0.023
	102131	U15085	Hs.1162	major histocompatibility complex; class II; DM beta	0.023
	101375	M13560	Hs.84298	CD74 entigen (invariant polypeptide of major histocompatibility	
	101070	MICOSO	11010-1210	complex: class it antigen-associated)	0.023
	100674	HG3033-HT3194		Spliceosomal Protein Sap 62	0.024
25	134365	R32377	Hs.82240	syntaxin 3A	0.027
23	132335	D60387	Hs.189885	ESTs	0.027
	110303	H37901	Hs.32706	ESTS	0.028
	131678	N59162	Hs 30542	ESTs	0.028
	116599	D80046	Hs.250879	ESTs	0.029
30	133769	M17733	Hs.75968	thymosin; beta 4; X chromosome	0.029
50	107904	AA026648	Hs.61389	ESTs	0.03
	129427	T80746	Hs.111334	ferritin; light polypeplide	0.03
	105987	AA406631	Hs.110299	milogen-activated protein kinase kinase 7	0.03
	131466	F03233	Hs.27189	ESTs	0.032
35	102859	X00274	Hs.76807	Human HLA-DR alpha-chain mRNA	0.032
33	134626	S82198	Hs.8709	caldecrin (serum calcium decreasing factor; elastase IV)	0.032
	134170	M63138	Hs.79572	cathepsin D (tysosomal aspartyl protease)	0.033
	131713	X57809	Hs.181125	Immunoglobulin lambda gene cluster	0.034
	100748	HG3517-HT3711	na. 101 123	Alpha-1-Antitryosin. 5' End	0.034
40	118769	N74498		ESTs	0.034
40	111734	R25375	Hs.126916	ESTs .	0.036
			Hs.85840	ESTs; Weakly similar to stac [H.saplens]	0.036
	109221	AA192755 AA480073	Hs.76719	U6 snRNA-associated Sm-like protein	0.036
	135281	AA401575	Hs.97757	ESTs -	0.037
45				v-ets avian erythroblastosis virus E26 oncogene related	0.037
43	119073	R32894	Hs.45514	Major Histocompatibility Complex, Class ti Beta W52	0.037
	100760	HG3576-HT3779	11-05		0.037
	101426	M19483	Hs.25	ATP synthase; H+ transpring; milochndrl F1 complex; beta polypept	0.038
	129568	AA428025	Hs.114360	transforming growth factor beta-stimulated protein TSC-22	0.036
50	130900	Z38468	Hs.21036	ESTs; Moderately similar to F25965_3 [H.sapiens]	0.039
30	133879	M13829	Hs.77183	v-raf murine sarcoma 3611 viral oncogene homolog 1	0.039
	100627	HG2702-HT2798		Serine/Threonine Kinase (Gb:Z25424)	0.039
	129424	M55593	Hs.111301	matrix metalloproteinase 2 (gelatinase A; 72kD gelatinase;	0.039
				72kD type IV collagenase)	0.039
	128652	AA621245	Hs.103147	ESTs; Weakly similar to similar to SP:YR40_BACSU [C.elegans]	0.039
55	129979	T72635	Hs.13956	ESTs	0.039
	133468	X03068	Hs.73931	major histocompatibility complex; class il; DQ beta 1	0.04
	102636	U67092		Human ataxia-telangiectasia locus protein (ATM) gene, exons	0.04
				1a, 1b, 2, 3 and 4, partial ods	0.04
60	129536	M33493	Hs.184504	tryptase; alpha	0.04
60	133599	M64788	Hs.75151	RAP1; GTPase activating protein 1	0.041

	102104	U12139		Human alpha1(XI) collagen (COL11A1) gene, 5' region and exon 1	0.041
	131340	AA478305	Hs.25817	Homo sapiens chromosome 19: cosmid R27216	0.041
	130446	X79510	Hs.155693	protein tyrosine phosphatase; non-receptor type 21	0.042
	101352	L77701	Hs.16297		0.042
5				COX17 (yeast) homolog; cytochrome c oxidase assembly protein	
,	122593	AA453310	Hs.128749	alpha-methylacyl-CoA racemase	0.042
	130181	R39552	Hs.151608	Homo sapiens clone 23622 mRNA sequence	0.042
	134071	Z14093	Hs.78950	branched chain keto acid dehydrogenase E1; alpha polypeptide	
				(maple syrup urine disease)	0.042
	108129	AA053252	Hs.185848		0.042
10	100125	MMUSSESE	FIS. 100040	ESTs; Weakly similar to II ALU SUBFAMILY J WARNING	
10				ENTRY !! [H.sapiens]	0.043
	130511	L32137	Hs.1584	cartilage oligomeric matrix protein (pseudoachondroplasia;	
				epiphyseal dysplasia 1; multiple)	0.043
	133336	AA291456	Hs.71190	ESTs	0.043
	132932	L02326	Hs.198118		
15				immunoglobulin fambda-like polypeptide 2	0.044
13	131880	AA047034	Hs.33818	RecQ protein-like 5	0.044
	130540	U35234	Hs.159534	protein tyrosine phosphatase; receptor type; S	0.044
	133467	AA258595	Hs.73931	major histocompatibility complex; class II; DQ beta 1	0.044
	101191	L20688	Hs.83666	Rho GDP dissociation Inhibitor (GDI) beta	0.044
	101860	M95610	Hs.37165		
20			HS.37 165	collagen; type IX; alpha 2	0.044
20	102799	U88898		Human endogenous retroviral H protease/integrase-derived ORF1	
				mRNA, complete cds, and putative envelope prot mRNA, partial cds	0.044
	107200	D20350	Hs.5628	ESTs	0.044
	101166	L14927	Hs.2099	Spocalin 1 (protein migrating faster than albumin; tear prealbumin)	0.044
		M54915		spotant i (protein ingland dater tran abunin, tear preabunin)	
25	134289		Hs.81170	pim-1 oncogene	0.044
25	135329	AA436026	Hs.98858	ESTs	0.044
	124950	T03786	Hs.151531	protein phosphatase 3 (tormerly 2B); catalytic subunit; beta isotom	
				(calcineurin A beta)	0.044
	102919	X12447	Hs.183760	aldolase A; fructose-bisphosphate	
			HS. 103/00		0.044
20	100574	HG2279-HT2375		Triosephosphate Isomerase	0.045
30	131286	AA450092	Hs.25300	Homo sapiens clones 24718 and 24825 mRNA sequence	0.045
	102675	U72512		Human B-cell receptor associated protein (hBAP) alternatively	
				spliced mRNA, partial 3'UTR	0.045
	131332	R50487	Hs.25717	ESTs	
					0.045
~ ~	101634	M57731	Hs.75765	GRO2 oncogene	0.046
35	113118	T47906	Hs.220512	ESTs	0.046
	124884	R77276	Hs.120911	ESTs	0.046
	130523	W76097	Hs.214507	ESTs	0.046
	110244				
		H26742	Hs.25367	ESTs; Weakly similar to ALR [H.saplens]	0.046
	131932	AA454980	Hs.25601	chromodomain helicase DNA binding protein 3	0.046
40	132509	H09751	Hs.5038	neuropathy target esterase	0.046
	133372	AA291139	Hs.72242	ESTs	0.046
	100817	HG4011-HT4804		Dystrophin-Associated Glycoprotein, 50 Kda, Alt. Splice 2	0.047
	106746	AA476436	11- 7004		
			Hs.7991	ESTs	0.047
	135401	L14813	Hs.169271	carboxyl ester lipase-like (bile salt-stimulated lipase-like)	0.047
45	130479	R44163	Hs.12457	Homo sapiens clone 23770 mRNA sequence	0.047
	102589	U62015	Hs.8867	cysteine-rich; angiogenic inducer; 61	0.047
	121521	AA412165	Hs.97358	EST	0.048
	135340	AA425137	Hs.99093	Homo sapiens chromosome 19; cosmid R28379	0.048
	132336	AA342422	Hs.45073	ESTs	0.048
50	115368	AA282133	Hs.88960	ESTs; Weakly similar to similar to collagen [C.elegans]	0.048
	101278	L38487	Hs.110849	estrogen-related receptor alpha	0.048
	103284	X80200	Hs.8375	TNF receptor-essociated factor 4	0.048
			F10.0075		
	100564	HG2239-HT2324		Potassium Channel Protein (Gb:Z11585)	0.048
	133132	Z40883	Hs.65588	ESTs; Weakly similar to dJ393P12.2 [H.sapiens]	0.048
55	121811	AA424535	Hs.98416	ESTs	0.046
	129613	AA279481	Hs.238831	ESTs; Weakly similar to collagen alpha 1(XVIII) chain [M.musculus]	0.049
	132468	\$79854	Hs.49322	deiodinase; iodothyronine; type III	
					0.049
	120111	W95841	Hs.136031	ESTs	0.049
	103668	Z83741	Hs.248174	H2A histone family; member M	0.049
60	130386	F10874	Hs.234249	mitogen-activated protein kinase 8 Interacting protein 1	0.049
	104275	C02170	Hs.39387	ESTs; Weakly smir to weak smirity to ribosomal prot L14 [C.elegans]	0.049
	106305	AA436146	Hs.12828	ESTS	
					0.05
	116431	AA609878	Hs.55289	ESTs; Weakly smir to 110 KD CELL MEMBRANE GLYCOPROTEIN [H.sapiens]	0.813
	120339	AA206465	Hs.258470	EST	0.05
65	114427	AA017063		ESTs; Highly similar to Miz-1 protein [H.saplens]	0.05
	118821	N79070	Hs.94789	ESTs	0.05
	118979	N93798	Hs.43666		
				protein tyrosine phosphatase type IVA; member 3	0.05
	107495	W78776	Hs.90375	ESTs	0.051
	120240	Z41732	Hs.66049	ESTs	0.051

		744600	11- 40400	ESTs	0.051
	114331	Z41309	Hs.12400		0.052
	130947	R40037	Hs.21506	ESTs	
	129242	W81679	Hs.5174	ribosomal protein S17	0.052
	131413	AA482390	Hs.26510	ESTs; Modily smlr to vacuolar prot sorting homolog r-vps33b [R.norvegicus]	0.052
5	112304	R54798	Hs.26239	ESTs	0.052
,	101416	M17254	Hs.45514	v-ets avian erythrobiastosis virus E26 oncogene related	0.052
					0.052
	131201	AA426304	Hs.24174	ESTs	
	101054	K02405	Hs.73933	Human MHC class II HLA-DQ-beta mRNA (DR7 DQw2); complete cds	0.052
	101306	L41143	Hs.232069	T-cell leukemia translocation altered gene	0.053
10	129311	T55087		vb45c08.r1 Stratagene fetal spleen (#937205) Homo sapiens cDNA	
10	120011	100007		done IMAGE:74126 5', mRNA sequence.	0.053
					0.053
	129942	U95301	Hs.144442	phospholipase A2; group X	
	119210	R93340	Hs.92995	ESTs	0.053
	101046	K01160		Accession not listed in Genbank	0.053
15	114096	Z38266	Hs.12770	Homo sapiens PAC clone DJ0777023 from 7p14-p15	0.053
13	110171	H19964	Hs.31709	ESTs	0.053
					0.053
	101004	J04101	Hs.248109	v-ets avian erythroblastosis virus E26 oncogene homolog 1	
	129715	N58479	Hs.12126	ESTs; Weakly similar to LR8 [H.saplons]	0.053
	101581	M34996	Hs.198253	major histocompatibility complex; class II; DQ alpha 1	0.053
20	113285	T66830	Hs.182712	ESTs	0.053
20		AA569531	Hs.162859	ESTs	0.054
	127537		HS. 102008		0.054
	100613	HG3995-HT4265		Cpg-Enriched Dna, Clone S19	
	101841	M93107	Hs.76893	3-hydroxybutyrate dehydrogenase (heart; mitochondrial)	0.054
	135053	R77159	Hs.93678	ESTs	0.054
25	101419	M17886	Hs.177592	ribosomal protein; large; P1	0.054
23			Hs.47622	ESTs	0.055
	119724	W69468	HS.47022		0.055
	102673	U72509		Human alternatively spliced B8 (B7) mRNA, partial sequence	
	129877	AA248589	Hs.13094	ESTs; Weakly similar to ORF YGR101w [S.cerevisiae]	0.055
	114788	AA156737	Hs.103904	EST	0.055
30	123812	AA620607	Hs.111591	ESTs	0.055
50				ESTs	0.055
	117669	N39237	Hs.44977		
	123782	AA610111	Hs.162695	EST	0.055
	102395	U41767	Hs.92208	a disintegrin and metalloproteinase domain 15 (metargidin)	0.055
	133795	M12529	Hs. 169401	apolipoprotein E	0.055
35	123193	AA489228	Hs.136956	ESTs	0.056
22					0.056
	132595	AA253369	Hs.155742	glyoxylate reductase/hydroxypyruvate reductase	
	104161	AA456471	Hs.7724	KIAA0963 protein	0.056
	115330	AA281145	Hs.88827	ESTs	0.056
	112693	T08000	Hs. 194684	bessoon (presynaptic cytomatrix protein)	0.056
40	133475	L29217	Hs.73987	CDC-like kinase 3	0.056
40					0.056
	128699	K03207	Hs.103972	proline-rich protein BatNI subfamily 4	
	102940	X13956	Hs.24998	Hu 12S RNA induced by poly(rl); poly(rC) and Newcastle disease virus	0.058
	131299	AA431464	Hs.25426	ESTs; Weakly similar to unknown [H.sapiens]	0.057
	102495	U51240	Hs.79356	Lysosomal-associated multispanning membrane protein-5	0.057
45	129594	R70379	Hs.115396	Human germline IgD chain gene; C-region; C-delta-1 domain	0.057
43					0.057
	118593	N69020	Hs.207689	EST	
	126702	U54602	Hs.2785	keratin 17	0.057
	124386	N27368	Hs.212414	sema domain: immunoglobulin domain (lg); short basic domain;	
				secreted; (semaphorin) 3E	0.057
50	130538	M20786	Hs.159509	alpha-2-plasmin inhibitor	0.057
50			Hs.22920		0.057
	114299	Z40782		similar to S68401 (cattle) glucose induced gene	
	115604	AA400378	Hs.49391	ESTs	0.057
	106052	AA416947	Hs.6382	ESTs: Highly similar to KIAA0612 protein [H.saplens]	0.057
	131730	U05681	Hs.31210	B-call CLL/lymphoma 3	0.057
55				ESTs; Modly smir to putative seven pass transmembrane prot [H.sapiens]	0.058
22	131285	AA479498	Hs.25274	ES 15; Modify Shift to pulsawe seven pass dansholinated proclatical proclatical	0.058
	129705	X78706	Hs.12068	carnifine acetyltransferase	
	123175	AA489010	Hs.178400	ESTs	0.058
	103592	Z30644	Hs.123059	chloride channel Kb	0.058
	118198	N5947B	Hs.48396	ESTs; Moderately similar to turnor necrosis factor-alpha	
60	110100		. 10.40000	-Induced protein B12 [H.sapiens]	0.058
OU				nicuceu proteiti duz (m.saptetis)	0.058
	104886	AA053348	Hs.144626	growth differentiation factor 11	0.058
	104250	AF000575	Hs.105928	leukocyte immunoglobulin-like receptor; subfamily B (with TM	
				and ITIM domains); member 3	0.058
	113301	T67452	Hs.13104	EST	0.058
65				ECT - Highly smirts must about hotograph 2A DD gamma galamit fill samburd	0.058
65	110441	H50302	Hs.19845	ESTs; Highly smlr to prot phosphatase 2A BR gamma subunit (H.saplens)	
	125297	Z39215	Hs.159409	ESTs	0.058
	135258	AA292423	Hs.97272	ESTs; Weakly similar to dJ281H8.2 [H.sapiens]	0.058
	130633	T92363	Hs.178703	ESTs	0.058
	112006	R42607	Hs.22241	hypothetical protein	0.058
	112006	n4200/	13.22241	нурошновом рюком	0.000

	130805	U12194	Hs.170238	sodium channel; voltage-gated; type I; beta polypeptide	0.058
	134907	D80002	Hs.178292	KIAA0180 protein	0.058
	132619	AA404565	Hs.53447	ESTs; Moderately similar to kinesin light chain 1 [M.musculus]	0.058
_	135115	N35489	Hs.94653	neurochondrin	0.058
5	100531	HG1872-HT1907		Major Histocompat/billty Complex, Dg	0.058
	124530	N62256	Hs.102727	EST	0.058
	119960	W87533	Hs.32699	ESTs; Moderately similar to LIV-1 protein [H.saplans]	0.058
	132793	AA478999	Hs.56966	KIAA0906 protein	0.058
10	101076	L04270	Hs.1116	lymphotoxin beta receptor (TNFR superfamily; member 3	0.058
10	130355	N92934	Hs.17409	cysteine-rich protein 1 (intestinal)	0.058
	134458 105904	AA192614 AA401452	Hs.83577 Hs.32060	cysteine and glycine-rich protein 3 (cardiac LIM protein) ESTs	0.059
	132878	AA026793	Hs.58679	ESTs; Wealdy similar to 4F2/CD98 light chain [M.musculus]	0.059
	121828	AA425166	Hs.98497	ESTs	0.059
15	133418	U76366	Hs.172727	Treacher Collins-Franceschetti syndrome 1	0.059
13	129317	N46244	Hs.110373	ESTs	0.059
	130153	D85815	Hs.15114	ras homolog gene family; member D	0.059
	124403	N31745	Hs.102493	ESTs	0.059
	127683	AA668123	Hs.134170	ESTs	0.059
20	129814	W20070	Hs.168625	KIAA0979 protein	0.059
	131770	D59682	Hs.31833	ESTs	0.06
	117557	N33920	Hs.44532	diublquitin	0.06
	103522	Y10514		H.sapiens mRNA for CD152 protein	0.06
	120029	W91960	Hs.250640	sequence-specific single-stranded-DNA-binding protein	0.06
25	102135	U15460	Hs.41691	activating transcription factor B	0.06
	123617	AA609183	Hs.181131	ESTs	0.06
	112136	R46100	Hs.9739	ESTs	0.061
	133725	V00563	Hs.179543	Immunoglobulin mu	0.061
20	102069	U09196	Hs.82520	Hu 1.1 kb mRNA upregitd in retinoic acid treated HL-60 neutrophilic cells	0.061
30	106555	AA455000	Hs.16725	ESTS	0.061
	123269	AA491226	Hs.105280	ESTs; Weakly similar to dJ963K23.2 [H.sapiens]	0.061
	109068	AA166837	Hs.72620	DKFZP434I114 prolein	0.061
	129399	AA263028	Hs.111076	malate dehydrogenase 2; NAD (mitochondrial)	0.061
35	129375 135271	W79850 AA397763	Hs.11081 Hs.97562	ESTs; Weakly similar to HPBRII-7 protein [H.sapiens] ESTs	0.061
33	132958	W90398	Hs.6147	KIAA1075 protein	0.061
	129364	AA477106	Hs.110757	DNA segment on chromosome 21 (unique) 2056 expressed sequence	0.061
	123427	AA598548	Hs.112471	ESTs	0.061
	105236	AA219179	Hs.19105	translocase of Inner mitochondrial membrane 17 (yeast) homolog B	0.061
40	101012	J04444	Hs.697	cytochrome c-1	0.062
	134791	L18983	Hs.89655	protein tyrosine phosphatase; receptor type; N	0.062
	133700	K01396	Hs.75621	protease inhibitor 1 (anti-elestase); alpha-1-antitrypsin	0.062
	123887	AA621065	Hs.112943	ESTs	0.062
	129363	H05704	Hs.110746	H sapiens HCR (a-helix coiled-coil rod homologue) mRNA; complete cds	0.062
45	105719	AA291644	Hs.36793	ESTS	0.062
	124226	H62396	Hs.190266	ESTS	0.062
	117437	N27645		yw5e3.s1 Weizmenn Olfactory Epithelium H sapiens cDNA clone	
				IMAGE:255676 3' smir to contains L1.13 L1 repetitive element;, mRNA seq	0.062
<b>CO</b>	132741	AA394133	Hs.55898	ESTs; Highly similar to OASIS protein [M.musculus]	0.062
50	134437	M26041	Hs.198253	major histocompatibility complex; class II; DQ alpha 1	0.062
	107664 120844	AA010594 AA349417	Hs.5326 Hs.96917	ESTs; Moderately similar to plm-1 protein [H.saplens] ESTs	0.062
	101574	M34182	Hs.158029	protein kinase; cAMP-dependent; catalytic; gamma	0.062
	131219	C00476	Hs.24395	small inducible cytokine subfamily B (Cys-X-Cys); member 14 (BRAK)	0.062
55	103495	Y09022	Hs.153591	Not56 (D. melanogaster)-like protein	0.062
55	129607	AA404594	Hs.11607	ESTS	0.062
	106467	AA450040	Hs.154162	ADP-ribosylation factor-like 2	0.062
	128841	T16358	Hs.106443	ESTs	0.062
	100515	HG1723-HT1729	1101100110	Macrophage Scavenger Receptor, All, Splice 2	0.062
60	119332	T54095		ESTs; Weakly similar to II ALU SUBFAMILY J WARNING ENTRY !! [H.saplens]	0,062
	134518	AA171939	Hs.23413	ESTs	0.062
	135012	X73608	Hs.93029	sparo/osteonectin; owov and kazal-like domains proteoglycan (testican)	0.063
	103575	Z26256		H.saplens isoform 1 gene for L-type calcium channel, exon 1	0.063
	115514	AA297739	Hs.55609	ESTs; Weakly similar to ISOLEUCYL-TRNA SYNTHETASE;	
65				CYTOPLASMIC [H.sapiens]	0.063
	103996	AA321355		EST2393 Bone marrow Homo sapiens cDNA 5' end, mRNA sequence	0.063
	110505	H55992	Hs.20495	DKFZP434F011 protein	0.063
	133912	X62744	Hs.77522 Hs.180255	major histocompatibility complex; class II; DM alpha	0.063
	129581	M33600	18.180200	major histocompatibility complex; class II; DR beta 1	0.003
				197	

	130139	R38280	Hs.150922	BCS1 (yeest homolog)-like	0.064
	105817	AA397825	Hs.5307	synaptopodin	0.064
	134658	AA410617	Hs.178009	ESTs	0.064
_	100306	D50495	Hs.80598	trenscription elongation fector A (SII); 2	0.064
5	100277	D42053	Hs.75890	site-1 protease (subtilisin-like; sterol-regulated; cleaves sterol regulatory	
				element binding proteins)	0.064
	133116	D61259	Hs.6529	ESTs	0.064
	134909	AA521488	Hs.90998	KIAA0128 protein	0.064
10	130319	X74794	Hs.154443	minichromosome maintenance deficient (S. cerevisiae) 4	0.064
10	132057	AA102469	Hs.173484	ESTs (1907/004) Harmonday - DNA	0.064
	108334	AA070473		zm7c8.s1 Stratagene neuroepithelium (#937231) Homo sapiens cDNA	0.064
	400700	FARRE	LI- 40070	clone IMAGE:5399 3', mRNA sequence KIAA0422 protein	0.064
	129763	F10815	Hs.12373 Hs.94617	ESTs: Weakly similar to predicted using Genefinder (C.elegans)	0.064
15	135112 122289	T67464 AA436856	Hs.98910	ESTs; Weakly similar to predicted using denember (C.elegans)  ESTs	0.064
13	133082	AA457129	Hs.6455	RuyB (E coli homolog)-like 2	0.064
	113213	T58607	HS.D400	ya94a02.s1 Stratagene placenta (#937225) Homo sapiens cDNA clone	0.004
	110210	130007		IMAGE:69290 3', mRNA sequence.	0.065
	106228	AA429290	Hs.17719	ESTs	0.065
20	130192	Y12661	Hs.171014	VGF nerve growth factor inductible	0.065
	104894	AA054087	Hs.18858	phospholipase A2; group IVC (cytosolic; calcium-independent)	0.065
	103508	Y10141		H.saplens DAT1 gene, partial, VNTR	0.065
	128474	U40671	Hs.100299	ligase III: DNA: ATP-dependent	0.065
	134012	AA417821	Hs.237924	ESTs; Highly similar to CGI-69 protein [H.sapiens]	0.065
25	134536	AA457735	Hs.850	IMP (Inosine monophosphale) dehydrogenase 1	0.065
	111714	R23146	Hs.23466	ESTs	0.065
	110521	H57060	Hs.108268	ESTs	0.065
	103282	X80198	Hs.77628	steroidogenic acute regulatory protein related	0.065
	113921	W80730	Hs.28355	ESTs	0.065
30	129331	N93465	Hs.110453	ESTs; Highly similar to CGI-38 protein [H.saplens]	0.065
	111318	N74597	Hs.180535	ESTs; Weakly similar to mitogen inducible gene mig-2 (H.sapiens)	0.065
	135138	AA036794	Hs.95196	ESTs; Weakly similar to T20B12.3 [C.elegans]	0.065
	107289	T10792	Hs.172098	ESTs	0.065
35	121405	AA406083	Hs.98007	ESTs ESTs	0.065
23	124965	T16275	Hs.106359 Hs.174481	ESIS ESTs	0.066
	106595 100106	AA456933 AF015910	FIS.1/4401	Homo sapiens unknown protein mRNA, partial cds	0.066
	134715	AA282757	Hs.89040	prepronociceptin	0.066
	135367	AA480109	Hs.9963	TYRO protein tyrosine kinase binding protein	0.066
40	111533	R08548	Hs.251651	EST	0.066
	128509	R53109	Hs.247362	dimethylarginine dimethylaminohydrolase 2	0.068
	101030	J05037	Hs.76751	serine dehydratase	0.066
	102753	U80226		Human gamma-aminobutyric acid transaminase mRNA, partial cds	0.067
	126991	R31652	Hs.821	bigiycan	0.067
45	109583	F02322	Hs.26135	ESTs	0.067
	119241	T12559	Hs.221382	ESTs	0.067
	130569	AA156597	Hs.256441	EST; Moderately similar to CGI-136 protein [H.sapiens]	0.067
	112926	T10316	Hs.4302	ESTs	0.067
~~	120495	AA256073	Hs.190628	ESTs	0.067
50	130931	AA278412	Hs.21346	ESTs; Weakly similar to F42C5.7 gene product [C.elegans]	0.067
	129982	M87789	Hs.140	Immunoglobulin gamma 3 (Gm marker)	0.067
	133832	H03387	Hs.241305	estrogen-responsive B box protein	0.067
	1 10697	H93721	Hs.20798	ESTs	0.067
55	121183	AA400138	Hs.97703	ESTs Wiskott-Aidrich syndrome (ecezema-thrombocytopenia)	0.067
33	130953	U12707 U24183	Hs.2157 Hs.75160	phosphofructokinase: muscle	0.067
	102218 114181	Z39079	Hs.8021	KIAA1058 protein	0.067
	116581	D51287	Hs.82148	ribosomai protein S12	0.067
	132498	T87708	Hs.50098	ESTs	0.068
60	103788	AA096014	Hs.9527	ESTs: Highly similar to HSPC013 [H.saplens]	0.068
00	102459	U48936	110.0027	Human amforde-sensitive epitheliel sodium channel gamme subunit mRNA,	0.068
	400070	Dinon	Hs.77225	5' end, partial ods	0.068
	100373	D79999 AA203321	Hs.77225 Hs.151696	ADP-ribosyltransferase (NAD+; poly (ADP-ribose) polymerase)-like 1 DKFZP727G051 protein	0.068
65	132717 128863	AA203321 D87462	Hs.101696 Hs.106674	BRCA1 associated protein-1 (ubiquitin carboxy-terminal hydrolase)	0.068
0.5	115193	AA262029	Hs.88218	ESTs	0.068
	124558	N66046	Hs.141605	ESTs	0.069
	117225	N20392	Hs.42846	ESTs	0.069
	110665	H83380	Hs.32757	ESTs	0.069

	132905	U70663	Hs.182965	Kruppel-like factor 4 (qut)	0.069
	105778	AA348910	Hs.153299	DOM-3 (C. elegans) homolog Z	0.069
	134770	R72079	Hs.89575	CD798 antigen (immunoglobulin-associated beta)	0.069
	123097	AA485869	Hs.105671	ESTs	0.069
5	100750	HG3523-HT4899		Proto-Oncogene C-Myc, Alt. Splice 3, Orf 114	0.069
-	125091	T91518		ye20f05.s1 Stratagene lung (#937210) H sapiens cDNA clone IMAGE:	
				3' similar to contains Alu repetitive element; contains MER 12 repetitive element;	
				mRNA sequence.	0.069
	100756	HG3565-HT3768		Zinc Finger Protein (Gb:M88357)	0.069
10	113483	T87768	Hs.16439	ESTs	0.069
	101119	L09708	Hs.2253	complement component 2	0.069
	102288	U31628	Hs.12503	interleukin 15 receptor; alpha	0.07
	135349	D83174	Hs.9930	collagen-binding protein 2 (colligen 2)	0.07
	100991	J03764	Hs.82085	plasminogen activator inhibitor; type I	0.07
15	133675	AA443720	Hs.7551	ESTs; Weakly similar to T25G3.1 [C.elegans]	0.07
	105422	AA251014	Hs.12210	ESTs	0.07
	102932	X13334	Hs.75827	CD14 antigen	0.07
	119147	R58878	Hs.65739	ESTs	0.07
	104900	AA055048	Hs.180481	ESTs; Weakly similar to ACROSIN PRECURSOR [H.sapiens]	0.07
20	133185	AA481404	Hs.6686	ESTs	0.07
	115496	AA290674	Hs.71819	eukaryotic translation initiation factor 4E binding protein 1	0.07
	121005	AA398332	Hs.97613	ESTs	0.07
	124869	R69038	Hs.28728	ESTs; Weakly similar to F55A12.9 [C.elegans]	0.071
0.5	129154	N23673	Hs.108969	mannosidase; alpha; class 2B; member 1	0.071
25	112161	R48295		ESTs; Wkly smlr to I! ALU SUBFAMILY J WARNING ENTRY II [H.sapiens]	0.071
	125251	W87486	Hs.141464	ESTs	0,077
	134298	J00116	Hs.81343	collagen; type il; alpha 1 (primary osteoarthritis; spondyloep(physeal	0.071
			11 50000	dysplasia; congenital)	0.071
20	119745	W70264	Hs.58093 Hs.25489	ESTs ESTs	0.071
30	131306	AA232686		ESIS ESTs	0.071
	107776	AA018820	Hs.221147	ESTs; Widy smir to II ALU SUBFAMILY SX WARNING ENTRY II [H.sapiens]	0.071
	134271	AA199630 M85220	Hs.184456	Accession not listed in Genbank	0.071
	101798		Hs.99922	dopamine receptor D4	0.071
35	135402	\$76942 N74052	Hs.50424	EST	0.071
33	118742 131867	N/4052 N64856	Hs.3353	Homo sapiens done 24940 mRNA sequence	0.071
	102923	X12517	Hs.1063	small nuclear ribonucleoprotein polypeptide C	0.072
	100775	HG371-HT26388	110.1000	Mucin 1, Epithelial, Alt. Splice 9	0.072
	111020	N54361	Hs.185726	ESTs	0.072
40	134224	X80822	Hs.163593	ribosomal protein L18a	0.072
40	124059	F13673	Hs.99769	ESTs	0.072
	133972	AA160743	Hs.78019	Homo sapiens clone 24432 mRNA sequence	0.072
	129681	AA436009	Hs.178186	ESTs: Weakly similar to WASP-tamily protein [H.sapiens]	0.072
	103065	X58399	Hs.81221	Human L2-9 transcript of unrearranged immunoglobulin V(H)5 pseudogene	0.072
45	124966	T19271	Hs.155560	calnexin	0.072
-13	112270	R53021	Hs.203358	ESTs	0.072
	116704	F10183	Hs.66140	EST	0.072
	129890	M13699	Hs.111461	coruloplasmin (ferroxidase)	0.072
	127345	AA972008	Hs. 166253	ESTs; Highly similar to KIAA0478 protein [H.sapiens]	0.072
50	112436	R63090	Hs.28391	ESTs	0.072
	114531	AA053033	Hs.203330	ESTs	0.072
	135122	H99060	Hs.94814	ESTs	0.072
	103934	AA281338	Hs.134200	Homo sepiens mRNA; cDNA DKFZp564C186 (from clone DKFZp564C186)	0.072
	109363	AA215369	Hs.185764	ESTs; Weakly similar to hypothetical protein [H.sapiens]	0.072
55	112647	R83329	Hs.33403	ESTs	0.073
	127083	Z44079	Hs.91608	otoferlin	0.073
	133027	AA402624	Hs.63236	synuclein; gamma (breast cancer-specific protein 1)	0.073
	122036	AA432121	Hs.250986	EST	0.073
	110405	H47542	Hs.33962	ESTs	0.073
60	128697	AB002344	Hs.103915	KIAA0346 protein	0.073
	112221	R50380	Hs.25670	ESTs	0.073
	100478	HG1067-HT1067		Mucin (Gb:M22406)	0.073
	115598	AA400129	Hs.65735	EST <sub>3</sub>	0.073
	132491	AA227137	Hs.4984	KIAA0628 protein	0.073
65	101655	M60299		Human alpha-1 collagen type II gene, exons 1, 2 and 3	0.073
	106018	AA411887	Hs.34737	ESTs	0.073
	129683	W05348	Hs.158196	DKFZP434B103 protein	0.073
	134137	F10045	Hs.79347	KIAA0211 gene product	0.073
	114008	W89128	Hs.19872	ESTs	0.073